

STIC Search Report Biotech-Chem Library

STIC Database Tracking Number 171478

TO: Everett White Location: 5d24 / 5c18

Thursday, November 17, 2005

Art Unit: 1623

Phone: 571-272-0660

Serial Number: 10 / 810742

From: Jan Delaval

Location: Biotech-Chem Library

Remsen 1a51

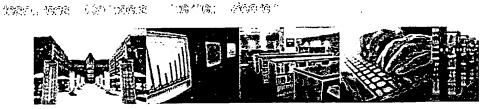
Phone: 571-272-2504

jan.delaval@uspto.gov

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-	







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Online Database Search Form

Search requests relating to published applications,	patent families,	and litigation car	n I
form and clicking on "Send."			

- Tech Center:
© TC 1600 € TC 1700 € TC 2100 € TC 2600 € TC 2800
C TC 2900 O TC 3600 O TC 3700 O Law Lib O Other
Your Contact Information: * indicates mandatory information.
Your Name: Everett White
*Email Address: Everett.White@uspto.gov
(e.g., Susan.Smith@uspto.gov)
*Employee No.: 67057
*Art Unit/Org.: 1623
*Office Location: REM 5D24
*Phone No.: 571-272-0660
Mailbox No.:
*Case serial number: 10/810,742 If not related to a patent application, please enter NA here.
Class / Subclass(es) 536/20 and 514/55
Earliest Priority Filing Date: 3/25/2004
Format preferred for results:
Paper Diskette E-mail

Provide detailed information on your search topic:

- In your own words, describe in detail the concepts or subjects you want us to se
- Include synonyms, keywords, and acronyms. Define terms that have special me
- *For Chemical Structure Searches Only*
 - Include the elected species or structures, keywords, synonyms, acronyms, and
- *For Sequence Searches Only* Include all pertinent information (parent, child, divisional, or issued patent numb serial number.
- *For Foreign Patent Family Searches Only* Include the country name and patent number.

- Provide examples or give us relevant citations, authors, etc., if known.
- FAX or send the abstract, pertinent claims (not all of the claims), drawings, c
 EIC or branch library.

Enter your Search Topic Information below:

Please carry out a structure-search of the N-acylated chitinous polymer having the formula set forth in instant Claims 1-11. A data base search of the cross linked n-acylated-N,O-carboxyalkylchitosan and a pharmaceutical composition thereof of Claims 42-50 is also requested. A data base search of a N-acylated-N,O-carboxyalkylchitosan being used to treat cancer, a nervous system disorder, a urinary tract disorder, a gastrointesting tract disorder, a reproductive tract disorder and to prevent surgical adhesion in a subject as set forth in Claims 24-44

Special Instructions and Other Comments:

(For fastest service, let us know the best times to contact you, in case the searcher no search.)

is further requested. A copy of the claims is enclosed

An inventor search (see Bib Data Sheet, enclosed) is als or requested.

I can be reach from about 11:00am to 5:00pm daily.



Submit comments and suggestions to Kristin Vajs

To report technical pro

If you cannot access a file because of a missing or non-working plugin, please contact the Help Desk at 2-9000 (Alexandria) or

305-9000 (Crystal City) for installation assistance.

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Last modified 11/14/2005 17:21:51

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=> d his
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(FILE 'HOME' ENTERED AT 13:42:09 ON 17 NOV 2005)
                SET COST OFF
     FILE 'REGISTRY' ENTERED AT 13:42:15 ON 17 NOV 2005
                E CHITIN/CN
L1
              1 S E3
L2
            315 S 1398-61-4/CRN
                E CHITIN
L3
           3200 S E3
L4
            924 S L3 NOT E4-E13, E16
L5
            608 S L4 NOT L1, L2
L6
             90 S L5 NOT SQL/FA
L7
             89 S L6 NOT DNA
L8
             11 S L2 AND N
L9
              5 S L8 AND C2H4O3
L10
              5 S L9 AND CARBOXYMETHYL ETHER
L11
              1 S 52519-63-8
L12
              8 S L1-L3 AND N AND O
L13
              4 S L12 NOT SQL/FA
     FILE 'HCAPLUS' ENTERED AT 13:50:01 ON 17 NOV 2005
L14
            375 S L11
           8628 S L1
L15
             12 S L15 (L) N(L)O
L16
                SEL AN 1 12
L17
              2 S L16 AND E1-E4
                SEL RN
     FILE 'REGISTRY' ENTERED AT 13:54:51 ON 17 NOV 2005
             16 S E5-E20
L18
L19
              2 S L18 AND L1-L3
L20
             14 S L18 NOT L19
L21
              1 S L20 AND C10H17NO8
     FILE 'HCAPLUS' ENTERED AT 13:57:06 ON 17 NOV 2005
L22
              6 S L21
                SEL AN 3-6
L23
              4 S L22 AND E21-E28
L24
              5 S L17, L23
              1 S US20050214255/PN OR (US2004-810742? OR WO2005-US10103)/AP,PRN
L25
                E ELSON C/AU
L26
            158 S E3-E8, E18, E19
                E KYDONIEUS A/AU
L27
            149 S E3-E10
                E HENDERSON S/AU
L28
             64 S E3, E10
                E HENDERSON SUE/AU
L29
              6 S E5, E9, E10
                E CHITOGEN/PA, CS
L30
             12 S E5-E12
L31
              4 S L26-L29 AND CHITIN
L32
              1 S L30 AND CHITIN
L33
              4 S L31, L32
L34
              2 S L33 NOT (48 OR 61)/SC, SX
L35
              2 S L33 NOT L34
                 SEL RN L34
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L36
             18 S E1-E18
L37
              4 S 1404-00-8 OR 56124-62-0 OR 89-57-6 OR 23214-92-8
L38
              2 S L36 AND (CHITIN OR L1-L3)
L39
              4 S L36 AND CHITOSAN
     FILE 'HCAPLUS' ENTERED AT 14:06:45 ON 17 NOV 2005
L40
          23166 S L38, L39
L41
             10 S L26-L30 AND L40
L42
              6 S L41 NOT L33
L43
              5 S L42 NOT 44/SC
L44
          28096 S L2, L3
L45
              0 S L26-L30 AND L44 NOT L41,L33
     FILE 'REGISTRY' ENTERED AT 14:09:24 ON 17 NOV 2005
L46
           1830 S CHITOSAN
     FILE 'HCAPLUS' ENTERED AT 14:09:32 ON 17 NOV 2005
L47
          19914 S L46
     FILE 'REGISTRY' ENTERED AT 14:09:45 ON 17 NOV 2005
L48
              1 S L39 AND 1/NC
L49
            894 S 9012-76-4/CRN
     FILE 'HCAPLUS' ENTERED AT 14:09:55 ON 17 NOV 2005
L50
           2267 S L49
L51
             21 S L26-L30 AND L47, L50
L52
             11 S L51 NOT L33, L41
             18 S L34, L43, L52
L54
             18 S L53 AND L14-L17, L22-L35, L40-L45, L47, L50-L53
L55
             17 S L54 AND N O
L56
             18 S L54 AND ?CHITOSAN?
L57
             3 S L54 AND ?CHITIN?
L58
             18 S L54-L57
L59
             17 S L58 AND ?CARBOXY?
L60
             18 S L58, L59
                SEL RN 18
     FILE 'REGISTRY' ENTERED AT 14:14:16 ON 17 NOV 2005
L61
             21 S E19-E39
L62
              3 S 865532-59-8 OR 865533-35-3 OR 865533-54-6
L63
              2 S L61 AND C5H9NO4
L64
              1 S L63 AND CHITOSAN
L65
              2 S L61 AND C6H8O7
L66
              1 S L65 AND CHITOSAN
L67
              1 S SUCCINIC ACID/CN
L68
           6216 S 110-15-6/CRN
                E C4H4O3/MF
L69
             43 S E3 AND OC4/ES
L70
              6 S L69 AND 2 5 NOT (14C# OR 13C# OR 11C# OR (D OR T)/ELS)
L71
              5 S L70 NOT DIOL
L72
              1 S L71 NOT (LABELED OR 5 HYDROXY)
L73
           1881 S 108-30-5/CRN
L74
              3 S L2, L3 AND L68, L73
                E C4H2O3/MF
L75
             16 S E3 AND OC4/ES
L76
              3 S L75 AND 2 5 NOT (14C# OR 13C# OR 11C# OR (D OR T)/ELS OR LABE
L77
              1 S 108-31-6
          24151 S 108-31-6/CRN
L78
L79
              0 S L2, L3 AND L78
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FILE 'HCAPLUS' ENTERED AT 14:24:27 ON 17 NOV 2005
L80
              2 S L62, L64, L66
L81
             23 S L17, L24, L25, L34, L43, L60, L80
L82
             23 S L81 AND L14-L17, L22-L35, L40-L45, L47, L50-L60, L80-L81
L83
             18 S L82 AND N O
L84
             23 S L82 AND (?CHITOSAN? OR ?CHITIN? OR ?CARBOXY? OR ?ACYL?)
L85
              1 S L84 AND L37
L86
              6 S L84 AND (5 AMINOSALICYLIC ACID OR DOXORUBICIN OR ?PEPTIDE? OR
L87
             10 S L84 AND NOCC
L88
             23 S L84-L87
L89
              4 S L14 AND L37
L90
             34 S L14 AND (5 AMINOSALICYLIC ACID OR DOXORUBICIN OR ?PEPTIDE? OR
L91
             35 S L89, L90
L92
              0 S L22 AND L37
L93
              0 S L22 AND (5 AMINOSALICYLIC ACID OR DOXORUBICIN OR ?PEPTIDE? OR
L94
             41 S L22, L91
L95
             40 S L94 AND (PD<=20040325 OR PRD<=20040325 OR AD<=20040325)
L96
             41 S L94, L95
                SEL AN 2 14 16 18 21 22 33 36 37
L97
             32 S L96 NOT E1-E18
     FILE 'REGISTRY' ENTERED AT 14:40:04 ON 17 NOV 2005
L98
              1 S DIVINYL SULFONE/CN
     FILE 'HCAPLUS' ENTERED AT 14:40:10 ON 17 NOV 2005
           1071 S L98 OR DIVINYLSULFONE OR DIVINYLSULPHONE OR DIVINYL() (SULFON
L99
L100
             21 S L99 AND L14, L15, L22, L40, L44, L47, L50
L101
              0 S L100 AND L97
L102
             33 S L25, L97
              1 S L102 AND L99, L100
L103
L104
             33 S L102,L103 AND L14-L17,L22-L35,L40-L45,L47,L50-L60,L80-L97,L99
                SEL HIT RN
     FILE 'REGISTRY' ENTERED AT 14:42:36 ON 17 NOV 2005
L105
             31 S E19-E49
L106
             25 S L105 AND (CHITOSAN OR CHITIN OR L2 OR L49)
L107
              9 S L106 AND N
              4 S L107 AND 1/NC
L108
              9 S L107, L108
L109
L110
              8 S L109 NOT C16H36N
              6 S L105 NOT L106
L111
L112
              1 S L111 AND PMS/CI
L113
              9 S L110, L112
L114
              5 S L111 NOT L113
     FILE 'HCAPLUS' ENTERED AT 14:45:47 ON 17 NOV 2005
L115
             33 S L113 AND L104
```

=> fil reg

FILE 'REGISTRY' ENTERED AT 14:46:12 ON 17 NOV 2005 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2005 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 16 NOV 2005 HIGHEST RN 868209-27-2 DICTIONARY FILE UPDATES: 16 NOV 2005 HIGHEST RN 868209-27-2

New CAS Information Use Policies, enter HELP USAGETERMS for details.

TSCA INFORMATION NOW CURRENT THROUGH JULY 14, 2005

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******************* * The CA roles and document type information have been removed from * * the IDE default display format and the ED field has been added, * effective March 20, 2005. A new display format, IDERL, is now * available and contains the CA role and document type information. * ******************

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REGISTRY includes numerically searchable data for experimental and predicted properties as well as tags indicating availability of experimental property data in the original document. For information on property searching in REGISTRY, refer to:

http://www.cas.org/ONLINE/UG/regprops.html

=> d l113 ide can tot

L113 ANSWER 1 OF 9 REGISTRY COPYRIGHT 2005 ACS on STN

865533-54-6 REGISTRY

ED Entered STN: 19 Oct 2005

CN Chitosan, N-(carboxymethyl)-N-[(2Z)-3-carboxy-2(or 3)-methyl-1-oxo-2-propenyl], carboxymethyl ether (9CI) (CA INDEX NAME)

ΜF C2 H4 O3 . x Unspecified

PCT Manual registration

SR

CA, CAPLUS, TOXCENTER, USPATFULL LC STN Files:

CM 1

865533-51-3 CRN CMF Unspecified

CCI PMS, MAN

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

CM 2

CRN 79-14-1 CMF C2 H4 O3

0 HO- C- CH2- OH

> 1 REFERENCES IN FILE CA (1907 TO DATE) 1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

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REFERENCE
          1: 143:353333
L113 ANSWER 2 OF 9 REGISTRY COPYRIGHT 2005 ACS on STN
RN
     865533-35-3 REGISTRY
ED
     Entered STN: 19 Oct 2005
     Chitosan, N-(carboxymethyl)-N-(4-carboxy-1-oxobutyl), carboxymethyl
CN
     ether (9CI) (CA INDEX NAME)
     C2 H4 O3 . x Unspecified
MF
PCT
    Manual registration
SR
     CA
LC
     STN Files:
                 CA, CAPLUS, TOXCENTER, USPATFULL
     CM
          1
         865533-33-1
     CRN
     CMF
         Unspecified
     CCI PMS, MAN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
     CM
          2
     CRN 79-14-1
     CMF C2 H4 O3
   0
HO-C-CH2-OH
               1 REFERENCES IN FILE CA (1907 TO DATE)
               1 REFERENCES IN FILE CAPLUS (1907 TO DATE)
REFERENCE
            1: 143:353333
L113 ANSWER 3 OF 9 REGISTRY COPYRIGHT 2005 ACS on STN
     865532-59-8 REGISTRY
ΕD
     Entered STN: 19 Oct 2005
CN
     Chitosan, N-(carboxymethyl)-N-(3-carboxy-1-oxopropyl), carboxymethyl
     ether (9CI) (CA INDEX NAME)
     C2 H4 O3 . x Unspecified
MF
    Manual registration
PCT
SR
LC
     STN Files:
                CA, CAPLUS, TOXCENTER, USPATFULL
     CM
     CRN
         865532-22-5
     CMF
         Unspecified
     CCI PMS, MAN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
     CM
          2
     CRN 79-14-1
     CMF C2 H4 O3
```

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О
||
НО- С- СН<sub>2</sub>- ОН
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1 REFERENCES IN FILE CA (1907 TO DATE)
1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 143:353333

L113 ANSWER 4 OF 9 REGISTRY COPYRIGHT 2005 ACS on STN

RN **83512-85-0** REGISTRY

ED Entered STN: 16 Nov 1984

CN Chitosan, N-(carboxymethyl) (9CI) (CA INDEX NAME)

OTHER NAMES:

CN Carboxymethylchitosan

CN N-Carboxymethylchitosan

MF Unspecified

CI PMS, COM, MAN

PCT Manual registration

LC STN Files: AGRICOLA, BIOBUSINESS, BIOSIS, CA, CANCERLIT, CAPLUS, CASREACT, CHEMCATS, CHEMLIST, CIN, CSCHEM, MEDLINE, PROMT, TOXCENTER, USPAT2, USPATFULL

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

353 REFERENCES IN FILE CA (1907 TO DATE)

48 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

356 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 143:374068

REFERENCE 2: 143:372878

REFERENCE 3: 143:365768

REFERENCE 4: 143:362030

REFERENCE 5: 143:348976

REFERENCE 6: 143:348654

REFERENCE 7: 143:341015

REFERENCE 8: 143:339391

REFERENCE 9: 143:332632

REFERENCE 10: 143:321142

L113 ANSWER 5 OF 9 REGISTRY COPYRIGHT 2005 ACS on STN

RN **78809-92-4** REGISTRY

ED Entered STN: 16 Nov 1984

CN Chitosan, N-(3-carboxy-1-oxopropyl) (9CI) (CA INDEX NAME)

OTHER NAMES:

CN Chitosan succinamate

CN Chitosan succinamide

```
white - 10 / 810742
CN
     Chitosan succinoyl amide
CN
    N-Succinylchitosan
MF
     Unspecified
CI
     PMS, COM, MAN
PCT
    Manual registration
LC
     STN Files:
                  BIOBUSINESS, BIOSIS, CA, CAPLUS, TOXCENTER, USPAT2, USPATFULL
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
              66 REFERENCES IN FILE CA (1907 TO DATE)
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              67 REFERENCES IN FILE CAPLUS (1907 TO DATE)
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            1: 143:253612
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            2:
               143:103223
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            3:
                143:78377
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                142:294296
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            5:
                142:242623
REFERENCE
                141:415826
REFERENCE
            7:
                141:337254
REFERENCE
            8:
                141:59792
REFERENCE
            9:
               141:28269
REFERENCE 10: 140:349983
L113 ANSWER 6 OF 9 REGISTRY COPYRIGHT 2005 ACS on STN
     57216-53-2 REGISTRY
ΕD
     Entered STN: 16 Nov 1984
     D-Glucose, 2-(acetylamino)-6-O-(carboxymethyl)-2-deoxy-, homopolymer (9CI)
CN
     (CA INDEX NAME)
```

(CA INDEX NAME)
OTHER NAMES:

CN Poly(N-acetyl-6-O-carboxymethyl-D-glucosamine)

FS STEREOSEARCH MF (C10 H17 N O8)x

CI PMS

PCT Polyamide, Polyamide formed, Polyester, Polyester formed, Polyether LC STN Files: CA, CAPLUS, IFICDB, IFIPAT, IFIUDB, TOXCENTER, USPATFULL

CM 1

CRN 57216-52-1 CMF C10 H17 N O8

Absolute stereochemistry.

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6 REFERENCES IN FILE CA (1907 TO DATE)
6 REFERENCES IN FILE CAPLUS (1907 TO DATE)
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REFERENCE 1: 112:223132

REFERENCE 2: 110:156586

REFERENCE 3: 94:180713

REFERENCE 4: 88:197685

REFERENCE 5: 88:126373

REFERENCE 6: 84:35314

L113 ANSWER 7 OF 9 REGISTRY COPYRIGHT 2005 ACS on STN

RN **52519-63-8** REGISTRY

ED Entered STN: 16 Nov 1984

CN Chitin, carboxymethyl ether (9CI) (CA INDEX NAME)

OTHER NAMES:

CN Carboxymethylchitin

CN N-Acetyl-O-carboxymethylchitosan

CN O-Carboxymethylchitin

DR 196412-80-3, 199943-94-7

MF C2 H4 O3 . x Unspecified

CI COM

PCT Manual registration, Polyother, Polyother only

LC STN Files: AGRICOLA, BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CANCERLIT, CAPLUS, CSCHEM, EMBASE, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, TOXCENTER, USPAT2, USPATFULL

CM 1

CRN 1398-61-4

CMF Unspecified

CCI PMS, MAN

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

CM 2

CRN 79-14-1

CMF C2 H4 O3

О || НО- С- СН₂- ОН

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

373 REFERENCES IN FILE CA (1907 TO DATE)

62 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

375 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 143:374068

REFERENCE 2: 143:292328

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REFERENCE
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            4: 143:254088
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                143:228311
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                143:138494
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            7:
                143:104071
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            8:
                143:103223
REFERENCE
            9:
                143:79650
REFERENCE 10: 143:61377
L113 ANSWER 8 OF 9 REGISTRY COPYRIGHT 2005 ACS on STN
     9012-76-4 REGISTRY
RN
     Entered STN: 16 Nov 1984
ΕD
CN
     Chitosan (8CI, 9CI) (CA INDEX NAME)
OTHER NAMES:
CN
     100D-VL
CN
     Amidan
CN
     BC 10
ÇN
     BC 10 (polysaccharide)
CN
     Biopolymer L 112
CN
     C 60M
CN
     Chicol
CN
     Chirosan 100
     Chitan, N-acetyl-
CN
CN
     Chitech
CN
     Chitin, N-deacetyl-
CN
     Chitoclear
CN
     Chitoclear 400
CN
     Chitoclear TM 1111
CN
     Chitoclear TM 588
CN
     Chitofos
CN
     Chitolaze
CN
     Chitopearl 3510
CN
     Chitopearl AL 10
CN
     Chitopearl BC 3000
CN
     Chitopearl BCW 2500
CN
     Chitopearl BCW 3000
CN
     Chitopearl BCW 3500
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     Chitopearl BCW 3505
CN
     Chitopearl BCW 3507
CN
     Chitopearl K 20
CN
     Chitosan 10B
CN
     Chitosan 500
CN
     Chitosan CLH
CN
     Chitosan EL
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     Chitosan F
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     Chitosan FL
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     Chitosan H
CN
     Chitosan LL
CN
     Chitosan LL 80
CN
     Chitosan LLWP
CN
     Chitosan M
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CN

Chitosan MP

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CN
     Chitosan PSH
CN
     Chitosan VL
CN
     Chitosan WL-M
CN
     Chitosol
CN
     Chitosom
CN
     Crystan LA-S
     CTA 1 Lactic Acid
CN
CN
     CTA 4
     DAC 50
CN
     DAC 70
CN
CN
     Daichitosan 100DVL
CN
     Daichitosan DVL
CN
     Daichitosan L
CN
     Daichitosan P-VL
CN
     Daichitosan VL
CN
     Daichitosan VLA
CN
     Daichitosan W 10
CN
     Deacetylchitin
CN
     Flonac N
CN
     FM 200 (chitosan)
CN
     K 5 (chitosan)
     Kimitsu Chitosan F
CN
CN
     Kimitsu Chitosan F 2
CN
     Kimitsu Chitosan F 2P
CN
     Kimitsu Chitosan H
CN
     Kimitsu Chitosan L
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     Kimitsu Chitosan LL
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     Kimitsu Chitosan LLW
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     Kimitsu Chitosan M
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     Kimitsu Chitosan MP
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MF
     Unspecified
CI
     PMS, COM, MAN
PCT
     Manual registration, Polyother, Polyother only
LC
     STN Files: ADISNEWS, AGRICOLA, ANABSTR, AQUIRE, BIOBUSINESS, BIOSIS,
       BIOTECHNO, CA, CABA, CANCERLIT, CAPLUS, CASREACT, CBNB, CEN, CHEMCATS,
       CHEMLIST, CIN, CSCHEM, CSNB, DDFU, DIOGENES, DRUGU, EMBASE, HSDB*,
       IFICDB, IFIPAT, IFIUDB, IMSDRUGNEWS, IMSRESEARCH, IPA, MEDLINE,
       NAPRALERT, PHAR, PIRA, PROMT, RTECS*, SCISEARCH, TOXCENTER, TULSA, USAN,
       USPAT2, USPATFULL, VTB
         (*File contains numerically searchable property data)
     Other Sources:
                      NDSL**, TSCA**, WHO
         (**Enter CHEMLIST File for up-to-date regulatory information)
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
**PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT**
           17392 REFERENCES IN FILE CA (1907 TO DATE)
```

jan delaval - 17 november 2005

2917 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA 17478 REFERENCES IN FILE CAPLUS (1907 TO DATE)

```
REFERENCE
            1: 143:397460
REFERENCE
             2: 143:397408
REFERENCE
             3: 143:393149
             4: 143:393144
REFERENCE
REFERENCE
             5: 143:393128
REFERENCE
             6: 143:393127
REFERENCE
            7: 143:393048
REFERENCE
             8: 143:393043
REFERENCE
            9: 143:393004
REFERENCE 10: 143:392960
L113 ANSWER 9 OF 9 REGISTRY COPYRIGHT 2005 ACS on STN
     1398-61-4 REGISTRY
RN
     Entered STN: 16 Nov 1984
F.D
CN
     Chitin (8CI, 9CI) (CA INDEX NAME)
OTHER NAMES:
CN
     α-Chitin
CN
     β-Chitin
CN
     γ-Chitin
CN
     Chitan, N-acetyl-
CN
     Chitin Tc-L
CN
     Clandosan
CN
     Kimica Chitin F
CN
     Kimitsu Chitin
CN
     North Chitin CG 2
CN
     Regitex FA
CN
     SEC 1
DR
     9043-70-3, 191802-95-6
MF
     Unspecified
CI
     PMS, COM, MAN
PCT
     Manual registration, Polyother, Polyother only
LC
     STN Files: AGRICOLA, ANABSTR, BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CABA,
       CANCERLIT, CAPLUS, CASREACT, CBNB, CHEMCATS, CHEMLIST, CIN, CSCHEM,
       CSNB, DDFU, DRUGU, EMBASE, HSDB*, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MRCK*, NAPRALERT, PROMT, RTECS*, TOXCENTER, USPAT2, USPATFULL, VETU, VTB
          (*File contains numerically searchable property data)
     Other Sources:
                       EINECS**, NDSL**, TSCA**
          (**Enter CHEMLIST File for up-to-date regulatory information)
```

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

8606 REFERENCES IN FILE CA (1907 TO DATE) 1050 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA 8628 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 143:393172

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REFERENCE
            2: 143:392545
REFERENCE
            3: 143:389263
REFERENCE
            4: 143:388812
REFERENCE
               143:388811
            5:
REFERENCE
            6: 143:385848
REFERENCE
            7: 143:385360
REFERENCE
            8:
               143:383201
REFERENCE
            9: 143:383033
REFERENCE 10: 143:382914
=> d l114 ide can tot
L114 ANSWER 1 OF 5 REGISTRY COPYRIGHT 2005 ACS on STN
     56124-62-0 REGISTRY
ED
     Entered STN: 16 Nov 1984
CN
     Pentanoic acid, 2-[(2S,4S)-1,2,3,4,6,11-hexahydro-2,5,12-trihydroxy-7-
     methoxy-6,11-dioxo-4-[{2,3,6-trideoxy-3-[(trifluoroacetyl)amino}]-\alpha-L-
     lyxo-hexopyranosyl]oxy]-2-naphthacenyl]-2-oxoethyl ester (9CI) (CA INDEX
     NAME)
OTHER CA INDEX NAMES:
     Pentanoic acid, 2-[1,2,3,4,6,11-hexahydro-2,5,12-trihydroxy-7-methoxy-6,11-
     dioxo-4-[[2,3,6-trideoxy-3-[(trifluoroacetyl)amino]-\alpha-L-lyxo-
     hexopyranosyl]oxy]-2-naphthacenyl]-2-oxoethyl ester, (2S-cis)-
OTHER NAMES:
    AD 32
CN
CN
     Antibiotic AD 32
     N-Trifluoroacetyladriamycin 14-valerate
CN
     N-Trifluoroacetyldoxorubicin 14-valerate
CN
CN
     NSC 246131
CN
     Trifluoroacetyladriamycin 14-valerate
CN
     Valrubicin
CN
     Valstar
FS
     STEREOSEARCH
     136816-53-0
DR
MF
     C34 H36 F3 N O13
                ADISINSIGHT, ADISNEWS, BEILSTEIN*, BIOBUSINESS, BIOSIS,
LC
       BIOTECHNO, CA, CANCERLIT, CAPLUS, CASREACT, CHEMCATS, CIN, DDFU,
       DIOGENES, DRUGU, EMBASE, HSDB*, IFICDB, IFIPAT, IFIUDB, IMSDRUGNEWS,
       IMSPATENTS, IMSRESEARCH, IPA, MEDLINE, MRCK*, PHAR, PROMT, PROUSDDR, PS,
       RTECS*, SYNTHLINE, TOXCENTER, USAN, USPAT2, USPATFULL
         (*File contains numerically searchable property data)
```

Absolute stereochemistry.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

201 REFERENCES IN FILE CA (1907 TO DATE)

14 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

201 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 143:360087

REFERENCE 2: 143:353333

REFERENCE 3: 143:185899

REFERENCE 4: 143:166641

REFERENCE 5: 143:146730

REFERENCE 6: 143:120685

REFERENCE 7: 142:480782

REFERENCE 8: 142:457053

REFERENCE 9: 142:457052

REFERENCE 10: 142:441852

L114 ANSWER 2 OF 5 REGISTRY COPYRIGHT 2005 ACS on STN

RN **23214-92-8** REGISTRY

ED Entered STN: 16 Nov 1984

CN 5,12-Naphthacenedione, 10-[(3-amino-2,3,6-trideoxy-α-L-lyxo-hexopyranosyl)oxy]-7,8,9,10-tetrahydro-6,8,11-trihydroxy-8-(hydroxyacetyl)-1-methoxy-, (8S,10S)- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN 5,12-Naphthacenedione, 10-[(3-amino-2,3,6-trideoxy-α-L-lyxo-hexopyranosyl)oxy]-7,8,9,10-tetrahydro-6,8,11-trihydroxy-8-(hydroxyacetyl)-1-methoxy-, (8S-cis)-

OTHER NAMES:

CN 14-Hydroxydaunomycin

```
CN
     Biotransdox
CN
     Caelyx
CN
     Doxil
CN
     Doxorubicin
CN
     Evacet
CN
     Hydroxydaunomycin
CN
     NSC 123127
CN
     PK 2
CN
     Rubex
FS
     STEREOSEARCH
DR
     24385-08-8, 25311-50-6, 23257-17-2, 29042-30-6
ΜF
     C27 H29 N O11
CT
LC
     STN Files:
                  ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, AQUIRE, BEILSTEIN*,
       BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CABA, CANCERLIT, CAPLUS, CASREACT,
       CBNB, CEN, CHEMCATS, CHEMLIST, CIN, CSCHEM, CSNB, DDFU, DIOGENES, DRUGU,
       EMBASE, HSDB*, IFICDB, IFIPAT, IFIUDB, IMSCOSEARCH, IMSDRUGNEWS,
       IMSPATENTS, IMSRESEARCH, IPA, MEDLINE, MRCK*, MSDS-OHS, NAPRALERT,
       NIOSHTIC, PATDPASPC, PHAR, PROMT, PROUSDDR, PS, RTECS*, SCISEARCH,
       TOXCENTER, USAN, USPAT2, USPATFULL, VETU
         (*File contains numerically searchable property data)
     Other Sources:
                    DSL**, EINECS**, WHO
         (**Enter CHEMLIST File for up-to-date regulatory information)
```

Absolute stereochemistry.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

15391 REFERENCES IN FILE CA (1907 TO DATE) 1039 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA 15435 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 143:392785

REFERENCE 2: 143:392716

REFERENCE 3: 143:392670

REFERENCE 4: 143:387283

REFERENCE 5: 143:387282

REFERENCE 6: 143:384777

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REFERENCE
           7: 143:384770
REFERENCE
           8: 143:383854
REFERENCE
           9: 143:381737
REFERENCE 10: 143:380778
L114 ANSWER 3 OF 5 REGISTRY COPYRIGHT 2005 ACS on STN
RN
     1404-00-8 REGISTRY
ED
     Entered STN: 16 Nov 1984
CN
    Mitomycin (8CI, 9CI) (CA INDEX NAME)
     Unspecified
MF
CI
     COM, MAN
                 ADISNEWS, AGRICOLA, ANABSTR, BIOBUSINESS, BIOSIS, BIOTECHNO,
LC
     STN Files:
       CA, CABA, CAPLUS, CBNB, CEN, CIN, CSNB, DIOGENES, EMBASE, IMSCOSEARCH,
       MEDLINE, MRCK*, NIOSHTIC, PROMT, TOXCENTER, USPAT2, USPATFULL
         (*File contains numerically searchable property data)
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
             917 REFERENCES IN FILE CA (1907 TO DATE)
             128 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
             922 REFERENCES IN FILE CAPLUS (1907 TO DATE)
           1: 143:373313
REFERENCE
REFERENCE
            2:
               143:365654
REFERENCE
            3:
               143:353333
               143:321809
REFERENCE
            4:
REFERENCE
            5:
               143:312023
REFERENCE
            6:
               143:311993
REFERENCE
            7:
               143:292623
REFERENCE
            8:
               143:260361
REFERENCE
            9: 143:260354
REFERENCE 10: 143:245758
L114 ANSWER 4 OF 5 REGISTRY COPYRIGHT 2005 ACS on STN
     89-57-6 REGISTRY
RN
ED
     Entered STN: 16 Nov 1984
     Benzoic acid, 5-amino-2-hydroxy- (9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
     Salicylic acid, 5-amino- (8CI)
OTHER NAMES:
CN
     2-Hydroxy-5-aminobenzoic acid
CN
     3-Carboxy-4-hydroxyaniline
CN
     5-Amino-2-hydroxybenzoic acid
CN
     5-Aminosalicylic acid
CN
     5-ASA
CN
     Asacol
     Asacolitin
CN
```

CN

Asacolon

```
CN
     Canasa
     Claversal
CN
CN
     Fisalamine
CN
     Ipocol
CN
     Lixacol
CN
     m-Aminosalicylic acid
CN
     Mesacol
     Mesalamine
CN
CN
     Mesalazine
CN
     Mesasal
     NSC 38877
CN
CN
     Pentasa
CN
     Rowasa
CN
     Salofalk
CN
     Salozinal
FS
     3D CONCORD
DR
     61513-32-4
MF
     C7 H7 N O3
CI
     COM
                  ADISNEWS, AGRICOLA, ANABSTR, AQUIRE, BEILSTEIN*, BIOBUSINESS,
LC
     STN Files:
       BIOSIS, BIOTECHNO, CA, CANCERLIT, CAPLUS, CASREACT, CBNB, CEN, CHEMCATS,
       CHEMINFORMRX, CHEMLIST, CIN, CSCHEM, CSNB, DDFU, DIOGENES, DRUGU,
       EMBASE, GMELIN*, HODOC*, IFICDB, IFIPAT, IFIUDB, IMSCOSEARCH,
       IMSDRUGNEWS, IMSPATENTS, IMSRESEARCH, IPA, MEDLINE, MRCK*, MSDS-OHS,
       PHAR, PROMT, PS, RTECS*, SPECINFO, SYNTHLINE, TOXCENTER, ULIDAT, USAN,
       USPAT2, USPATFULL
         (*File contains numerically searchable property data)
                     DSL**, EINECS**, TSCA**, WHO
         (**Enter CHEMLIST File for up-to-date regulatory information)
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PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1852 REFERENCES IN FILE CA (1907 TO DATE)
97 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
1856 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 143:393095 REFERENCE 2: 143:392993 REFERENCE 3: 143:373078 REFERENCE 4: 143:360097 REFERENCE 5: 143:360095 REFERENCE 6: 143:359795 REFERENCE 7: 143:353333

REFERENCE 8: 143:338774

REFERENCE 9: 143:338769

REFERENCE 10: 143:304736

L114 ANSWER 5 OF 5 REGISTRY COPYRIGHT 2005 ACS on STN

RN **77-77-0** REGISTRY

ED Entered STN: 16 Nov 1984

CN Ethene, 1,1'-sulfonylbis- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Vinyl sulfone (6CI, 8CI)

OTHER NAMES:

CN Bis(ethenyl)sulfone

CN Divinyl sulfone

CN NSC 133793

CN NSC 18590

CN NSC 57304

FS 3D CONCORD

MF C4 H6 O2 S

CI COM

LC STN Files: AGRICOLA, BEILSTEIN*, BIOBUSINESS, BIOSIS, CA, CANCERLIT, CAOLD, CAPLUS, CASREACT, CBNB, CEN, CHEMCATS, CHEMINFORMRX, CHEMLIST, CIN, CSCHEM, DETHERM*, HODOC*, HSDB*, IFICDB, IFIPAT, IFIUDB, MEDLINE, NIOSHTIC, PIRA, PROMT, RTECS*, SPECINFO, SYNTHLINE, TOXCENTER, TULSA, USPAT2, USPATFULL

(*File contains numerically searchable property data)
Other Sources: EINECS**

(**Enter CHEMLIST File for up-to-date regulatory information)

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

756 REFERENCES IN FILE CA (1907 TO DATE)

88 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

756 REFERENCES IN FILE CAPLUS (1907 TO DATE)

44 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

REFERENCE 1: 143:389851

REFERENCE 2: 143:388551

REFERENCE 3: 143:388155

REFERENCE 4: 143:367713

REFERENCE 5: 143:353333

REFERENCE 6: 143:347207

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REFERENCE 7: 143:329274
REFERENCE 8: 143:326400
REFERENCE 9: 143:289413
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REFERENCE 10: 143:286442

=> fil hcaplus FILE 'HCAPLUS' ENTERED AT 14:46:31 ON 17 NOV 2005 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2005 AMERICAN CHEMICAL SOCIETY (ACS)

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FILE COVERS 1907 - 17 Nov 2005 VOL 143 ISS 21 FILE LAST UPDATED: 16 Nov 2005 (20051116/ED)

New CAS Information Use Policies, enter HELP USAGETERMS for details.

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> => d l115 bib abs hitrn retable tot

```
L115 ANSWER 1 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN
AN
     2005:1050467 HCAPLUS
DN
     143:353333
     N-acylated chitinous polymers and methods of use
TΙ
     thereof
IN
     Elson, Clive; Kydonieus, Agis; Henderson, Susan
     Elizabeth
PA
     Chitogenics, Inc., USA
SO
     U.S. Pat. Appl. Publ., 11 pp.
     CODEN: USXXCO
DТ
     Patent
LA
     English
FAN.CNT 1
     PATENT NO.
                         KIND
                                DATE
```

```
APPLICATION NO.
                                                                DATE
                                          -----
                               20050929
                                         US 2004-810742
PΙ
    US 2005214255
                        Α1
                                                                 20040325 <--
    WO 2005094278
                        A2
                               20051013
                                          WO 2005-US10103
                                                                 20050325 <--
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
            CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
            GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
            LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
            NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM,
            SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM,
        RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM,
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AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK,
             EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT,
             RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML,
             MR, NE, SN, TD, TG
PRAI US 2004-810742
                                20040325 <--
                          Α
AB
     The invention pertains to N-acetylated, N, O-
     carboxyalkylchitosans and methods for using the chitosans
     to treat disorders, such as cancer, nervous system disorders, urinary
     tract disorders, and reproductive tract disorders.
IT
     89-57-6, 5-Aminosalicylic acid
     1404-00-8, Mitomycin 23214-92-8, Doxorubicin
     56124-62-0, Valrubicin
     RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (N-acylated chitinous polymers and methods of use
        thereof)
ΙT
     77-77-0, Divinyl sulfone
     RL: RCT (Reactant); RACT (Reactant or reagent)
        (N-acylated chitinous polymers and methods of use
        thereof)
ΙT
     1398-61-4DP, Chitin, N-acylated analogs
     9012-76-4DP, Chitosan, N-acylated derivs.
     9012-76-4DP, Chitosan, N-acylated-N,
     O-carboxyalkyl derivs. 865532-59-8P
     865533-35-3P 865533-54-6P
     RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological
     study); PREP (Preparation); USES (Uses)
        (N-acylated chitinous polymers and methods of use
        thereof)
L115 ANSWER 2 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN
     2005:564589 HCAPLUS
     143:103223
DN
ΤI
     Polymeric micellar complexes and drug delivery vehicles thereof
ΙN
     Ignatious, Francis; Li, Yue Hu
     Smithkline Beecham Corporation, USA
PA
SO
     PCT Int. Appl., 24 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     English
FAN.CNT 1
     PATENT NO.
                                           APPLICATION NO.
                         KIND
                                DATE
                                                                  DATE
                         ____
                                            -----
PΙ
     WO 2005058376
                         A1
                                20050630
                                           WO 2004-US42768
                                                                   20041217 <--
             AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
             CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
             GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
             LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
             NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
             TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
         RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM,
             AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK,
             EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT,
             RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML,
             MR, NE, SN, TD, TG
PRAI US 2003-530142P
                         Ρ
                                20031217 <--
     US 2003-532045P
                          Ρ
                                20031222 <--
AB
     Disclosed are complexes of an amphiphilic copolymer, wherein the
     amphiphilic copolymer has benzoyl sulfonic acid groups on the hydrophobic
     segment of the copolymer. Poly(lactide-block-methoxypolyethylene glycol)
```

```
was functionalized with sulfobenzoic anhydride.
    1398-61-4, Chitin 9012-76-4, Chitosan 52519-63-8
TΤ
    , Carboxymethyl chitin 78809-92-4 83512-85-0,
    Carboxymethyl chitosan
    RL: POF (Polymer in formulation); THU (Therapeutic use); BIOL (Biological
    study); USES (Uses)
       (polymeric micellar complexes and drug delivery vehicles)
ፐጥ
    23214-92-8, Doxorubicin
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
       (polymeric micellar complexes and drug delivery vehicles)
RETABLE
  Referenced Author | Year | VOL | PG | Referenced Work | Referenced
       (RAU)
                   |(RPY)|(RVL)|(RPG)| (RWK)
|1984 |185 |1795 |Makromolekulare Chem|HCAPLUS
L115 ANSWER 3 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN
    2004:914821 HCAPLUS
DN
    142:162042
TI
    Method of preparing aqueous solution of chitin derivative and hyaluronic
    acid for cosmetic and medical purposes
ΙN
    Kim, Han Seok
PΑ
SO
    Repub. Korean Kongkae Taeho Kongbo, No pp. given
    CODEN: KRXXA7
DT
    Patent
T.A
    Korean
FAN.CNT 1
                   KIND DATE
    PATENT NO.
                                      APPLICATION NO.
    -----
                     ----
                             -----
                                        ------
                           20010928 KR 2001-50131 20010820 <--
    KR 2001088675 A
PRAI KR 2001-50131
                             20010820 <--
   A method of preparing a gel-like aqueous solution for cosmetic and medical
purposes
    by adding purified hyaluronic acid as a kind of mucopolysaccharide to
    carboxymethylchitin is provided. Whereby, the aqueous solution has excellent
    moisturizing action, tissue regenerating power and antibacterial activity
    and fine line wrinkles removing effect. The gelled aqueous solution is
prepared by
    mixing carboxymethyl chitin and 0.2 to 0.5% by weight of hyaluronic acid,
    optionally a thickening agent, vitamin, herb oil, antibiotics
    and other additives.
    52519-63-8, Carboxymethylchitin
    RL: COS (Cosmetic use); BIOL (Biological study); USES (Uses)
       (method of preparing aqueous solution of chitin derivative and hyaluronic
acid for
       cosmetic and medical purposes)
L115 ANSWER 4 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN
    2004:794550 HCAPLUS
DN
    141:282891
TI
    Hemostatic wound dressings and methods of making same
ΙN
    Looney, Dwayne Lee; Crilley, John; Guo, Jian Xin; Zhang, Guanghui;
    Pendharkar, Sanyog Manohar
PA
    Ethicon, Inc., USA
SO
    Eur. Pat. Appl., 50 pp.
    CODEN: EPXXDW
DT
    Patent
LA
   English
FAN.CNT 1
```

```
PATENT NO.
                                         APPLICATION NO. DATE
                         KIND
                                 DATE
                      ----
                         A1 20040929 EP 2003-254107
                                                                    -----
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     EP 1462123
PΙ
                                                                  20030627 <--
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
     US 2004193088
                        A1
                                 20040930
                                          US 2003-396224 20030325 <--
     CA 2433976
                         AA
     CA 2433976 AA 20040925 CA 2003-2433976 20030627 <--
CN 1531910 A 20040929 CN 2003-127787 20030627 <--
JP 2004290649 A2 20041021 JP 2003-185768 20030627 <--
BR 2003004169 A 20050517 BR 2003-4169 20030630 <--
US 2003-396224 A 20030325 <--
                                 20040925 CA 2003-2433976
                                                                    20030627 <--
PRAI US 2003-396224
     The present invention is directed to methods of making wound dressings
     that include the steps of (i) providing a solution of a water-soluble or
     water-swellable biocompatible polymer dissolved in a solvent for the
     polymer, (ii) providing a fabric substrate having a first surface and a
     second surface opposing the first surface, the fabric having properties
     effective for use as a hemostat and containing fibers prepared from a
     biocompatible polymer, (iii) contacting the fabric substrate with the
     polymer solution under conditions effective to distribute the polymer solution
     substantially homogeneously on the first and second surfaces and through
     the fabric substrate, (iv) transferring the fabric substrate having the
     polymer solution substantially homogeneously distributed there through to a
     lyophilization unit under conditions effective to maintain the homogeneous
     distribution on and throughout the substrate, and (v) lyophilizing the
     fabric having the polymer solution distributed there through, thereby
     providing a porous, polymeric matrix substantially homogeneously
     distributed on the first and second surfaces and through the fabric, the
     matrix being made-up of the lyophilized water-soluble or water-swellable
     polymer. For example, 1 g of hydroxyethyl cellulose (HEC) was dissolved
     in 99 g of water, and 10 g of the HEC solution was transferred into a
crystallization
     dish with a diameter of 10 cm. A piece of Surgicel Nu-Knit (absorbable
     hemostat, based on carboxylic-oxidized regenerated cellulose (CORC)),
     having a diameter of 9.8 cm (about 1.3 g) was placed on the HEC solution After
     soaking the fabric in the solution for 3 min, the wet fabric in the dish was
     then lyophilized overnight. A very flexible patch was formed. The patch
     achieved 100% hemostasis within 2 min in a porcine spleen incision model.
IΤ
     1398-61-4, Chitin 9012-76-4, Chitosan 52519-63-8
     , Carboxymethyl chitin
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (hemostatic dressings containing fabric substrate and porous, water-soluble
or
        water-swellable biocompatible polymeric matrix)
L115 ANSWER 5 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN
AN
     2004:586812 HCAPLUS
DN
     141:128434
ΤI
     Oxidative hair dye compositions containing polypeptides, chitin
     derivatives, or chitosan derivatives
     Chikakura, Yoshito; Nozaki, Kiyotada; Kato, Miyuki; Kono, Kenji; Miyamoto,
ΙN
     Kunihiro; Kitahara, Jiro; Nakata, Satoru
PA
     Nonogawa Shoji Ltd., Japan
     Jpn. Kokai Tokkyo Koho, 11 pp.
SO
     CODEN: JKXXAF
DT
     Patent
     Japanese
I.A
```

KIND DATE APPLICATION NO.

A2

DATE

20040722 JP 2002-374455 20021225 <--

FAN.CNT 1

PΙ

PATENT NO.

JP 2004203778

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PRAI JP 2002-374455 20021225 <--
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AB Oxidative hair dyes contain polypeptides (average mol. weight 200-30,000), chitin derivs., or chitosan derivs. as dyeing aids. The dyeability and color fastness to shampooing of 2-component oxidative hair dye compns. were significantly improved by addition of 0.1 weight% keratin hydrolyzate (average mol. weight 20,000) to the 1st component containing p-aminophenol, ammonium thioglycolate, etc.

IT 39280-86-9, Hydroxyethyl chitosan 52519-63-8,
 Carboxymethyl chitin 724422-43-9, Hydroxymethyl chitosan
 RL: COS (Cosmetic use); BIOL (Biological study); USES (Uses)
 (oxidative hair dye compns. containing polypeptides, chitin
 derivs., or chitosan derivs. as dyeing aids)

L115 ANSWER 6 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN

AN 2004:510784 HCAPLUS

DN 141:59786

- TI Hemostatic wound dressing made of a polysaccharide fabric and a polymer matrix
- IN Zhang, Guanghui; Pendharkar, Sanyog Manohar; Guo, Jian Xin; Looney, Dwayne Lee; Gorman, Anne Jessica

PA USA

SO U.S. Pat. Appl. Publ., 19 pp.

CODEN: USXXCO

DT Patent

- LA English
- FAN.CNT 7

	PA	rent	NO.			KIN)	DATE		I	APPL	ICAT	ION I	NO.		D2	ATE		
PI	US 2004120993 CA 2433977 EP 1430911			A1 AA A2		2004 2004 2004					20	00212 00306 00306	627	<					
	ΕP	1430 R:	911, AT,	BE,	CH,	A3 DE,		2004 ES,		GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,	
	CN	1509		SI,	LT,	LV, A	•	•	MK, 0707	•		TR, 003-			•		SK 0030	627	<
		2004 2003				A2 A			0722 0831			003- 003-				_	00306 00306		
PRAI		2004 2002				A1 A2			0603 0628			003-	7218	36		20	0031:	125	<
		2002 2002				A2 A2			1126 1126	<									
	US	2002 2002	-326	244		A2 A		2002	1126 1220		-								
	US	2003	-396	226		A2		2003	0325	<	-								

- AB The present invention is directed to hemostatic wound dressings containing a fabric made from biocompatible, aldehyde-modified polysaccharide fibers; and a porous, polymeric matrix made from a biocompatible, water-soluble or water-swellable polymer, dispersed at least partially through the fabric. The wound dressing further comprises a hemostatic agent, e.g., prothrombin, thrombin, fibrinogen, fibrin, fibronectin, heparinase, blood coagulation factors, tissue factor, batroxobin, ancrod, ecarin, etc. Methods of making such wound dressings and methods of providing hemostasis to a wound using the dressing are also described. For example, an aldehyde-modified regenerated cellulose fabric was soaked with a solution containing hydroxyethyl cellulose and thrombin and lyophilized to give a flexible patch. The patch achieved effective hemostasis in 73 s in a porcine splenic incision model with tamponade for 30 s.
- IT 1398-61-4D, Chitin, oxidized 9012-76-4D, Chitosan, oxidized 52519-63-8D, Carboxymethyl chitin, oxidized 83512-85-0D, Carboxymethylchitosan, oxidized

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (wound dressing containing aldehyde-modified polysaccharide fabric, polymer matrix and hemostatic)

L115 ANSWER 7 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN

AN 2004:510350 HCAPLUS

DN 141:59783

TI Bone repair materials and osteogenic cell culture base materials comprising peptide-grafted bioabsorbable polymers

IN Muramatsu, Kazuaki

PA Kyocera Corp., Japan

SO Jpn. Kokai Tokkyo Koho, 7 pp.

CODEN: JKXXAF
DT Patent

LA Japanese

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
		KIND	DAIL	AFFLICATION NO.	DATE
PΙ	JP 2004173995	A2	20040624	JP 2002-344739	20021127 <
PRAI	JP 2002-344739		20021127	<	

AB The bone repair materials and osteogenic cell culture base materials comprise bioabsorbable polymers having graft chains of **peptides** containing acidic amino acid-rich regions of bone marker proteins as motifs. An aqueous solution containing pepsin-treated type I collagen was

freeze-dried, and the resulting spongy sheet was irradiated with Ar plasma, immersed in an aqueous solution containing an osteocalcin **peptide** E-P-R-R-E-V-C-E-L-N-P-D-C-D-E, and freeze-dried to give a culture base material. Subcultured fibroblasts were cultured on the base materials in αMEM supplemented with 15% FBS, dexamethasone, ascorbic acid, and β -glycerophosphoric acid for 1 wk to give a material showing increased alkaline phosphatase activity, which was completely absorbed after 4-wk implantation into rabbit bone and enhanced the mineralization of bone.

IT 52519-63-8DP, Carboxymethyl chitin, graft polymers with osteonectin peptide

RL: SPN (Synthetic preparation); TEM (Technical or engineered material use); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(bone repair materials and osteogenic cell culture base materials comprising bioabsorbable polymers grafted with **peptides** containing bone marker protein motifs)

L115 ANSWER 8 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN

AN 2004:412562 HCAPLUS

DN 140:412378

TI Anti-adhesion compositions of polyacids and polyethers for reducing post-surgical pain

IN Schwartz, Herbert E.; Blackmore, John M.; Cortese, Stephanie M.; Oppelt,
William G.; DiZigera, Gere

PA USA

SO U.S. Pat. Appl. Publ., 86 pp., Cont.-in-part of U.S. Ser. No. 472,110. CODEN: USXXCO

DT Patent

LA English

FAN.CNT 6

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
ΡI	US 2004096422	A1	20040520	US 2003-666804	20030919 <
	US 5906997	Α	19990525	US 1997-877649	19970617 <
	US 6034140	A	20000307	US 1998-23097	19980213 <

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US 6869938
                          B1
                                20050322
                                            US 1999-472110
                                                                   19991227 <--
     WO 2005027852
                                            WO 2004-US30839
                          A2
                                20050331
                                                                   20040920 <--
     WO 2005027852
                          Α3
                                20051027
             AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
             CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
             GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
             LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
             NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
             TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
         RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM,
             AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK,
             EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE,
             SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE,
             SN, TD, TG
PRAI US 1997-877649
                          A3
                                19970617 <--
     US 1998-23097
                          Α2
                                19980213 <--
     US 1999-127571P
                          Ρ
                                19990402
                                         <--
     US 1999-472110
                          A2
                                19991227
                                         <--
     US 2003-666804
                          Α
                                20030919 <--
AΒ
     The present invention relates to improved methods for reducing pain and
     organ dysfunction using bioadhesive, bioresorbable, anti-adhesion compns.
     made of intermacromol. complexes of carboxyl-containing polysaccharides,
     polyethers, polyacids, polyalkylene oxides, multivalent cations and/or
     polycations. The polymers are associated with each other, and are then
     either dried into membranes or sponges, or are used as gels, fluids or
     microspheres. Compns. are useful in surgery to prevent the formation and
     reformation of post-surgical adhesions. The compns. are designed to
     breakdown in-vivo, and thus be removed from the body. Membranes are
     inserted during surgery either dry or optionally after conditioning in aqueous
     solns. Anti-adhesion, bioadhesive, bioresorptive, antithrombogenic and
     phys. properties of such membranes and gels can be varied as needed by
     carefully adjusting the pH and/or cation content of the polymer casting
     solns., polyacid composition, the polyalkylene oxide composition, or by
conditioning
     the membranes prior to surgical use. Membranes and gels can be used
     concurrently. Anti-adhesion compns. may also be used to lubricate tissues
     and/or medical instruments, and/or deliver drugs to the surgical site and
     release them locally. For example, an ionically crosslinked gel having 2%
     weight/volume solids ratio and 95% CM-cellulose/5% polyethylene oxide was
     prepared A dry, powdered mixture containing 9.5 g CMC and 0.5 g PEO was added
to 500
    mL water containing 3.2 mL of a 25.2% weight/volume solution of FeCl2.6H2O and
the
     solution was stirred at high speed until homogeneous. The osmolality was
     then adjusted to a physiol. acceptable value of about 300 mmol/kg by
     adding about 13 mL of a 30% weight/volume solution of NaCl and further mixing
the
     gel. The pH of the gel was adjusted to 7.0 by adding 1.7 N NH4OH.
     gel was sterilized in an autoclave for 15 min at 250°. Freeze
     drying of the gel provided iron-associated sponges.
ΙT
     1398-61-4, Chitin 52519-63-8, Carboxymethyl chitin
     83512-85-0, Carboxymethylchitosan
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (anti-adhesion compns. of polyacids and polyethers for reducing
       post-surgical pain)
L115 ANSWER 9 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN
ΑN
     2004:3477 HCAPLUS
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Hemostatic wound dressing comprising biocompatible polymeric fibers

DN

TI

140:65281

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Guo, Jian Xin; Looney, Dwayne Lee; Zhang, Guanghui; Gorman, Anne Jessica
IN
PA
     USA
     U.S. Pat. Appl. Publ., 17 pp.
SO
     CODEN: USXXCO
DT
     Patent
     English
LA
FAN.CNT 7
     PATENT NO.
                         KIND
                                DATE
                                           APPLICATION NO.
                                                                    DATE
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ΡI	US	2004	0018	79		A1	2004	0101	US	2002-	1860	21		2	0020	628	<
	US	2004	0053	50		A1	2004	0108	US	2003-	3962	26		2	0030	325	<
	CA	2433	968			AA	2003	1228	CA	2003-	2433	968		2	0030	627	<
	EΡ	1378	255			A2	2004	0107	EP	2003-	2541	14		2	0030	627	<- -
	EΡ	1378	255			А3	2004	0128									
		R:	AT,	BE,	CH,	DE,	DK, ES,	FR,	GB, GR	, IT,	LI,	LU,	NL,	SĖ,	MC,	PT,	
			ΙE,	SI,	LT,	LV,	FI, RO,							HU,	SK	·	
	CN	1533	751			Α	2004	1006	CN	2003-	1274	81		2	0030	627	<
	JP	2004	1601	82		A2	2004	0610	JP	2003-	1873	44		2	0030	630	<
	US	2004	1063	44		A1	2004	0603	US	2003-	7218	36		2	0031	125	<
PRAI	US	2002	-186	021		A2	2002	0628	<								
	US	2002	-304	472		A2	2002	1126	<								
	US	2002	-304	781		A2	2002	1126	<								
	US	2002	-305	040		A2	2002	1126	<								
	US	2002	-326	244		A2	2002	1220	<								
	US	2003	-396	226		A2	2003	0325	<								

AB The present invention is directed to wound dressings that contain a fabric made from biocompatible polymeric fibers and having flexibility, strength and porosity effective for use as a hemostat, and a porous, polymeric matrix prepared from a biocompatible, water-soluble or water-swellable polymer dispersed through the fabric; and to methods of making such wound dressings. For example, 1 g of hydroxyethyl cellulose (HEC) was dissolved in 99 g of water, and 10 g of the HEC solution was transferred into a crystallization

dish. A piece of Surgicel Nu-Knit absorbable hemostat, based on oxidized regenerated cellulose (ORC), having a diameter of 9.8 cm (about 1.3 g) was placed on the HEC solution in the crystallization dish. After soaking the fabric in

the solution for 3 min, the wet fabric was lyophilized to give a very flexible patch.

TΤ 1398-61-4, Chitin 9012-76-4, Chitosan 52519-63-8 , Carboxymethyl chitin 83512-85-0, Carboxymethyl chitosan RL: DEV (Device component use); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(hemostatic wound dressing comprising fabric made of biocompatible polymeric fibers and polymeric matrix)

- L115 ANSWER 10 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN
- 2003:790594 HCAPLUS ΑN
- DN 140:292561
- ΤI Development of degradable double-phase material promoting the regeneration of osteochondral defects in the knee joint
- ΑU Yoshihara, Yusuke; Mukai, Ken
- CS Biochemical Research Section, Bioceram Division, Kyocera Corporation, Japan
- SO Baiomateriaru (2003), 21(4), 305-310 CODEN: BAIOBS; ISSN: 1347-7080
- PΒ Nippon Baiomateriaru Gakkai
- DT Journal
- LA Japanese
- AB In the previous animal expts., the authors confirmed that

Carboxymethyl-chitin (CM-chitin) disappeared in rabbit tibial bone tissue with moderate inflammation, and did not obstruct the natural healing of bone defects. In this study, the authors manufactured CM-chitin by trial production for application to osteochondral defects and examined the ability in critical sized osteochondral defects of rabbits. Double phase materials (4.0 + 4.0 mm cylindrical construction), which are composed of CM-chitin and β -tricalcium phosphate (β -TCP) granules. were implanted into 5.0 + 5.0 mm osteochondral defects in the femoropatellar groove of young adult (4-mo) NZW rabbits. Defects filled with only β -TCP granules and defects without material were made as controls. Then, portions of the distal femur were harvested 2, 4 and 8 wk after implantation, decalcified and embedded in paraffin, and serial sections were cut at the center of the osteochondral defects. Sections were stained with toluidine blue solution, and microscopic examination was carried out. Two weeks later, in the case of double phase material implantation, the porous structure of CM-chitin had already disappeared, and the β -TCP granules were uniformly placed in newly formed tissue. The β -TCP granules gradually disappeared and active endochondral ossification was observed in the center of the defects. After eight weeks, ossification advanced further in the subchondral bone area and cartilaginous tissue remained on the upper side of the defect. On the other-hand, in the case of β -TCP granule implantation. ossification advanced in the subchondral bone area, but ultimately cartilaginous tissue unformed on the upper side of the defect. The result of this animal study indicates that the double phase material, which contains β -TCP granules in CM-chitin porous matrix, promotes new subchondral bone formation at early stage, and ultimately leads to the regeneration of cartilage-like tissue in the upper area of the osteochondral defect. 52519-63-8, Carboxymethyl-chitin

TT 52519-63-8, Carboxymethyl-chitin
RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL

(carboxymethyl-chitin- β -tricalcium phosphate double-phase material promoting the regeneration of osteochondral defects in knee joint)

L115 ANSWER 11 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN

(Biological study); USES (Uses)

AN 2003:218703 HCAPLUS

DN 139:296834

- TI Tissue response to a newly developed calcium phosphate cement containing succinic acid and carboxymethyl-chitin
- AU Yokoyama, Atsuro; Matsuno, Hironobu; Yamamoto, Satoru; Kawasaki, Takao; Kohgo, Takao; Uo, Motohiro; Watari, Fumio; Nakasu, Masanori
- CS Department of Oral Functional Science, Hokkaido University Graduate School of Dental Medicine, Sapporo, 060-8586, Japan
- SO Journal of Biomedical Materials Research, Part A (2003), 64A(3), 491-501

CODEN: JBMRCH

- PB John Wiley & Sons, Inc.
- DT Journal
- LA English

to

AB We developed a new calcium phosphate cement containing succinic acid and carboxymethyl-chitin in the liquid component. In this study, the biocompatibility and osteocond. of this new cement were investigated. After mixing, cement in putty form was implanted immediately between the periosteum and parietal bone and in the s.c. tissues of rats. In control cement, distilled water was used instead of the liquid component. In addition

histol. evaluations, analyses with x-ray diffraction and Fourier transform IR were performed for the s.c. implanted cements. Histol. examination showed slight **inflammation** around the new cement on the bone and in the s.c. tissue at 1 wk after surgery. At 2 wk, the cement was partially

bound to the parietal bone. The extent of the surface of the new cement directly in contact with the bone increased with time, and most of the undersurface of the new cement bound to the host parietal bone by 8 wk. Anal. by x-ray diffraction showed that the new cement in the s.c. tissue was transformed into hydroxyapatite by 8 wk. These results indicate that this new calcium phosphate cement is useful as a bone substitute material. 52519-63-8, Carboxymethyl-chitin

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (tissue response to a newly developed calcium phosphate cement containing succinic acid and carboxymethyl-chitin)

RETABLE

TΤ

RETABLE					
Referenced Author	Year	VOL	PG	Referenced Work	Referenced
(RAU)	(RPY)	(RVL)	(RPG)	(RWK)	File
=======================================	-+=====	+====	+=====	+======================================	
Bermudez, O	11994	15	160	J Mater Sci Mater Me	HCAPLUS
Brown, W	11986	1	1	US 4612053	HCAPLUS
Constantz, B	11995	1267	1796	Science	HCAPLUS
Donath, K	11987	113	120		MEDLINE
Frankenburg, E	1998	180-A	11112	J Bone and Joint Sur	
Friedman, C	1998	143	1428	J Biomed Mater Res (HCAPLUS
Fukase, Y	11990	169	1852	J Dent Res	HCAPLUS
Goodman, S	1990	24	517	J Biomed Mater Res	HCAPLUS
Hong, Y	11991	125	1485	J Biomed Mater Res	HCAPLUS
Hupp, J	1988	46	1538	J Oral Maxillofac Su	MEDLINE
Ikenaga, M	11998	40	1139	J Biomed Mater Res	HCAPLUS
Ishikawa, K	11995	16	1527	Biomaterials	HCAPLUS
Ishikawa, K	11995	6	1528	J Mater Sci Mater Me	HCAPLUS
Jarcho, M	11981	157	1259	Clin Orthop	HCAPLUS
Kent, J	11986	4 4	137	J Oral Maxilofac Sur	MEDLINE
Kurashina, K	11998	19	1701	Biomaterials	HCAPLUS
Kurashina, K	11995	6	1340	J Mater Sci Mater Me	
Miyamoto, Y	11997	37	1457	J Biomed Mater Res	HCAPLUS
Miyamoto, Y	11999	48	136	J Biomed Mater Res (HCAPLUS
Monma, H	11976	84	1209	J Ceram Soc Jpn	HCAPLUS
Monma, H	1997	15	124	J Jp Soc Biomaterial	HCAPLUS
Nakasu, M			Ì	Biomaterials; submit	
Sarkar, M	12001	58	1329	J Biomed Mater Res (HCAPLUS
Sjogren, U	11995	103	313	Eur J Oral Sci	MEDLINE
Takechi, M	11998	19	12057	Biomaterials	HCAPLUS
Wan, A	1996	17	1529	Biomaterials	HCAPLUS
Watanabe, K	1990	38	506	Chem Pharm Bull (Tok	HCAPLUS
Wittkampf, A	1988	46	1019	J Oral Maxillofac Su	MEDLINE
Yokoyama, A	11994	73	914	J Dent Res	
Yokoyama, A	11999	78	410	J Dent Res	
Yokoyama, A	•	44	9	J Jpn Prosthodont So	
Yuan, H	•		1283	-	HCAPLUS

L115 ANSWER 12 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN

AN 2002:107093 HCAPLUS

DN 136:156446

TI Mucoadhesive pharmaceutical composition comprising photochemotherapeutic agent

IN Klaveness, Jo; Hansson, Vidar; Godal, Aslak

PA Photocure ASA, Norway; Golding, Louise

SO PCT Int. Appl., 37 pp. CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

PATENT NO. KIND DATE APPLICATION NO. DATE

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WO 2002009690
PΙ
                         A2
                               20020207
                                         WO 2001-GB3338 20010725 <--
                               20020808
    WO 2002009690
                        A3
    WO 2002009690
                        C1
                               20040415
        W:
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
            CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
            GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
            LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT,
            RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US,
            UZ, VN, YU, ZA, ZW
        RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AM, AZ, BY, KG,
            KZ, MD, RU, TJ, TM, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR,
            IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN,
            GQ, GW, ML, MR, NE, SN, TD, TG
    CA 2417533
                        AΑ
                               20020207
                                         CA 2001-2417533
                                                                 20010725 <--
    EP 1311259
                                        EP 2001-984399
                         Α2
                               20030521
                                                                 20010725 <--
    EP 1311259
                        В1
                               20050615
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
            IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
    JP 2004505040
                        Т2
                               20040219
                                        JP 2002-515243
                                                                 20010725 <--
    AT 297723
                        E
                               20050715
                                         AT 2001-984399
                                                                 20010725 <--
    US 2004029855
                        A1
                               20040212
                                         US 2003-343211
                                                                20030805 <--
PRAI GB 2000-18527
                        Α
                               20000727
                                        <--
    WO 2001-GB3338
                        W
                               20010725 <--
OS
    MARPAT 136:156446
AΒ
    The present invention relates to a pharmaceutical composition for use as a
    medicament, preferably for the treatment or diagnosis of disorders or
    abnormalities of epithelial-lined surfaces, preferably mucosa-lined
    surfaces, comprising a photochemotherapeutic agent together with a
    mucoadhesive agent, optionally together with at least one surface
    penetration assisting agent and optionally with one or more chelating
    agents, and products and kits for performing the invention. A
    mucoadhesive composition contained hexyl 5-aminolevulinic acid hydrochloride
    1.5, liquid paraffin 2.0, and Orabase paste q.s. 100 q. The formulation
    showed improved mucoadhesive properties over the controls when applied to
    the mouth, tongue, and between the teeth of a man.
    9012-76-4, Chitosan 52519-63-8, Carboxymethyl chitin
    66267-50-3, Chitosan lactate 84563-76-8, Chitosan
    qlutamate
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (mucoadhesive pharmaceutical composition comprising photochemotherapeutic
       agent)
L115 ANSWER 13 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN
AN
    2002:6790 HCAPLUS
DN
    136:42523
ΤI
    Make-up material composition containing chitinous substance derivatives
ΙN
    Kwon, Sun Sang; Jang, Eui Seop; Song, Dong Hyuk; Yeom, Myung Hun; Moon,
    Chang Bae; Ahn, Soo Sun; Kim, Jin Han
PA
    Pacific Co., Ltd., S. Korea
    Repub. Korean Kongkae Taeho Kongbo, No pp. given
SO
    CODEN: KRXXA7
DT
    Patent
LA
    Korean
FAN.CNT 1
                               DATE APPLICATION NO. DATE
    PATENT NO.
                      KIND
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                                                                 _____
    KR 2000002616
PΙ
                               20000115
                                        KR 1998-23460
                                                                19980622 <--
PRAI KR 1998-23460
                               19980622 <--
    A skin preparation which is excellent in curing wounds and provides
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anti-inflammatory and moisturizing effects, comprises
carboxymethyl chitin. The carboxymethyl chitin is obtained by
carboxymethylation of chitin originated from cuttlefish bone.

IT 52519-63-8P, Carboxymethyl chitin

RL: COS (Cosmetic use); PAC (Pharmacological activity); PUR (Purification or recovery); BIOL (Biological study); PREP (Preparation); USES (Uses) (skin prepns. containing chitin ethers)

L115 ANSWER 14 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN

AN 2000:725477 HCAPLUS

DN 133:286502

- TI Compositions of polyacids and polyethers and methods for their use in reducing adhesions
- IN Schwartz, Herbert E.; Blackmore, John M.; Cortese, Stephanie M.; Oppelt, William G.
- PA Fziomed, Inc., USA
- SO PCT Int. Appl., 189 pp.

CODEN: PIXXD2

- DT Patent
- LA English

FAN.CNT 6

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PATENT NO.
                        KIND
                               DATE
                                          APPLICATION NO.
                                                                  DATE
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PΙ
    WO 2000059516
                         A1
                               20001012
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                                                                  20000323 <--
        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR,
            CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU,
            ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU,
            LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE,
            SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW
        RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE,
            DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF,
            CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
    US 6869938
                               20050322
                                           US 1999-472110
                                                                  19991227 <--
                         В1
    CA 2366880
                         AA
                               20001012
                                           CA 2000-2366880
                                                                  20000323 <--
                               20020227
                                           EP 2000-921450
    EP 1181023
                         Α1
                                                                  20000323 <--
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
            IE, SI, LT, LV, FI, RO
                         Т2
                               20031014
                                           JP 2000-609079
    JP 2003530136
                                                                  20000323 <--
    AU 778853
                         В2
                               20041223
                                          AU 2000-41770
                                                                  20000323 <--
                        Ρ
PRAI US 1999-127571P
                               19990402
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    US 1999-472110
                        Α
                               19991227
                                         <--
    US 1997-877649
                         A3
                               19970617
                                         <--
    US 1998-23097
                         A2
                               19980213
                                         <--
    WO 2000-US7963
                         W
                               20000323 <--
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AB The present invention relates to improved methods for making and using bioadhesive, bioresorbable, anti-adhesion compns. made of intermacromol. complexes of carboxyl-containing polysaccharides, polyethers, polyacids, polyalkylene oxides, multivalent cations and/or polycations. The polymers are associated with each other, and are then either dried into membranes or sponges, or are used as fluids or microspheres. Bioresorbable, bioadhesive, anti-adhesion compns. are useful in surgery to prevent the formation and reformation of post-surgical adhesions. The compns. are designed to breakdown in-vivo , and thus be removed from the body. Membranes are inserted during surgery either dry or optionally after conditioning in aqueous solns. The anti-adhesion, bioadhesive, bioresorptive, antithrombogenic and phys. properties of such membranes and gels can be varied as needed by carefully adjusting the pH and/or cation content of the polymer casting solns., polyacid composition, the polyalkylene oxide composition, or by conditioning the membranes prior to surgical use. Multi-layered membranes can be made and used to provide further control

over the phys. and biol. properties of antiadhesion membranes. Membranes and gels can be used concurrently. Antiadhesion compns. may also be used to lubricate tissues and/or medical instruments, and/or deliver drugs to the surgical site and release them locally. An examples was given for preparation of a neutral CM-cellulose-PEG membrane.

IT 1398-61-4, Chitin 52519-63-8, Carboxymethyl chitin

83512-85-0, Carboxymethyl chitosan

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (compns. of polyacids and polyethers and methods for their use in reducing adhesions)

RETABLE

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- L115 ANSWER 15 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN
- AN 1997:751590 HCAPLUS
- DN 128:46478
- TI Cell adhesion molecules and cancer metastasis
- AU Saiki, Ikuo
- CS Research Institute for Wakan-yaku, Toyama Medical and Pharmaceutical University, Toyama, 930-01, Japan
- SO Japanese Journal of Pharmacology (1997), 75(3), 215-242 CODEN: JJPAAZ; ISSN: 0021-5198
- PB Japanese Pharmacological Society
- DT Journal; General Review
- LA English
- AB A review with 138 refs. The adhesive interaction between tumor cells and host cells or the extracellular matrix plays a crucial role in metastasis formation. Therefore, understanding the mechanism controlling metastasis may assist in the development of anti-metastatic therapy. The authors have used synthetic or recombinant polypeptide analogs containing the Arg-Gly-Asp (RGD) sequence found in the functional domains of fibronectin, such as poly(RGD) or CH-271, to regulate the mechanisms involved in cell adhesion during the metastatic process. Poly(RGD) inhibited exptl. lung and liver metastasis effectively when coinjected i.v. with various types of tumors. In a model of spontaneous lung metastasis using the B16-BL6 melanoma, repeated administration of this polypeptide before or after surgical excision of the primary tumor resulted in a significant inhibition of tumor metastasis without affecting the growth of the primary tumor and substantially prolonged the survival time of mice. The mechanism responsible for the inhibition of tumor metastasis by the polypeptides is at least partly associated with the ability to interfere with cellular functions such as adhesiveness, motility and invasiveness in the process of metastasis. Combined treatment of the CH-271 fusion polypeptide and anticancer drugs, i.e., anti-adhesion therapy combined with chemotherapy, caused a marked inhibition of lung and liver metastasis of tumors as compared with either treatment alone or with the control. In contrast, the promotion of tumor cell interaction with immune cells via cell adhesion mols., which differs

from the anti-adhesive mechanism, may lead to the induction of anti-tumor immune responses and, consequently, to the inhibition of tumor metastasis. The transfection of the gene of the B7-1 adhesion mol. into tumor cells (B16-BL6 or K1735-M2 melanoma) resulted in the remarkable reduction of lung metastasis caused by the i.v. injection into mice. Immunization of B7-transfected tumor was effective as a tumor vaccine for preventing the metastasis of B7 neg. original tumor cells. Thus, the regulation of the adhesive interaction with tumor cells may provide a new and promising approach for the control and prevention of cancer metastasis.

R	E	ГΑ	R	Τ.	F.

RETABLE					
Referenced Author	Year	VOL	PG	Referenced Work	Referenced
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Bevilacqua, M	1987	84	19238	Proc Natl Acad Sci U	HCAPLUS
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Butcher, E	1991	167	11033	Cancer Res Cell	MEDLINE
Chan, B	1991	251	11600		HCAPLUS
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Fujii, H	1995	118			HCAPLUS
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Menter, D	11987				HCAPLUS
Mentzer, S	11985	135	19	J Immunol	HCAPLUS
Meromsky, L	1986	46	15270	Cancer Res	HCAPLUS
Miyake, M	1991	30	13328	Biochem	HCAPLUS
Mortarini, R	1991	47	551	Int J Cancer	HCAPLUS
Mould, A	1991	266	13579	J Biol Chem	HCAPLUS
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Yoneda, J
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L115 ANSWER 16 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN

AN 1997:317813 HCAPLUS

DN 126:288107

TI Antiinflammatory agents containing chitin derivatives

IN Tokura, Seiichi; Minami, Saburo; Tanioka, Shinichiro; Myazaki, Satoshi

PA San Fuaibu Kk, Japan

SO Jpn. Kokai Tokkyo Koho, 5 pp.

CODEN: JKXXAF

DT Patent

LA Japanese

FAN.CNT 1

2				
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI JP 09059164	A2	19970304	JP 1995-212003	19950821 <
PRAT .TP 1995-212003		19950821	<	

Antiinflammatory agents contain phosphated chitin, phosphated carboxymethylated chitin, or their salts as active ingredients. Powdered chitin (1.2 kg) was purified from 10 kg calamaries. Then, the powdered chitin (10 g) was dispersed in a mixture of DMF and urea and treated with H3PO4 and then with aqueous NaOH to give .apprx.12 g chitin phosphate Na salt (I). I (at 5.4 mg/kg i.v.) was effective in treatment of stomatitis in cats.

IT 52519-63-8DP, Carboxymethyl chitin, phosphates 72429-67-5P 99549-27-6P 189084-81-9P

(03) ref.

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RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); PNU (Preparation, unclassified); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (preparation of chitin phosphates as antiinflammatory agents)
ΙT
     1398-61-4P, Chitin
     RL: PUR (Purification or recovery); RCT (Reactant); PREP (Preparation);
     RACT (Reactant or reagent)
        (preparation of chitin phosphates as antiinflammatory agents)
L115 ANSWER 17 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN
     1996:380786 HCAPLUS
ΑN
DN
     125:95891
TΙ
     Synthesis of CM-chitin/doxorubicin conjugate attached through
     tetrapeptide spacer groups and release behavior of
     doxorubicin from itself in, vitro
ΑU
     Ouchi, Tatsuro; Nonomura, Koji; Hirai, Keiichi; Ohya, Yuichi
CS
     Faculty Engineering, Kansai University, Suita, 564, Japan
SO
     Chitin World, [Proceedings from the International Conference on Chitin and
     Chitosan], 6th, Gdynia, Pol., Aug. 16-19, 1994 (1994), 350-356.
     Editor(s): Karnicki, Zbigniew S. Publisher: Wirtschaftsverlag NW,
     Bremerhaven, Germany.
     CODEN: 62YQAK
DT
     Conference
LA
     English
AB
    Chitin is a non-toxic, biocompatible and biodegradable polysaccharide.
     6-O-carboxymethyl-chitin (CM-chitin) is a water-soluble chitin derivative In
     order to provide a water-soluble macromol. prodrug of doxorubicin
     (DXR) reducing the side-effects and exhibiting high antitumor activity,
     the fixation of DXRs to CM-chitin through covalent bonds was carried out.
     Two kinds of conjugate, the CM-chitin/Gly-Phe-Leu-Gly/DXR conjugate having
     lysosomally digestible tetrapeptide spacer groups and the
     CM-chitin/C5/DXR conjugate having pentamethylene spacer groups, were
     synthesized and the effect of the kind of spacer group species on the
     release behavior of DXR from the conjugates was investigated. The
     CM-chitin/Gly-Phe-Leu-Gly/DXR conjugate showed the specific fast release
     rate of DXR in the presence of the lysosomal enzyme (cathepsin B) at
     37° in vitro. On the contrary, the CM-chitin/C5/DXR conjugate did
     not show such a specific release behavior. Furthermore, the antitumor
     activities of these conjugates were investigated in vitro and in vivo.
TΤ
     1398-61-4, Chitin 23214-92-8D, Doxorubicin,
     spacer group conjugates with carboxymethylchitin 52519-63-8D,
     Carboxymethylchitin, spacer group conjugates with doxorubicin
     RL: RCT (Reactant); RACT (Reactant or reagent)
        (synthesis of carboxymethylchitin-doxorubicin conjugate
        attached through tetrapeptide spacer groups and release
       behavior of doxorubicin from itself)
L115 ANSWER 18 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN
ΑN
     1996:71305 HCAPLUS
DN
     124:126885
ΤI
     Skin cosmetics containing natural salt
ΙN
     Nakagawa, Momoki
PA
SO
     Jpn. Kokai Tokkyo Koho, 3 pp.
     CODEN: JKXXAF
DΤ
     Patent
LA
     Japanese
FAN.CNT 1
     PATENT NO.
                       KIND
                                DATE
                                           APPLICATION NO.
                                                                  DATE
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PΙ
    JP 07304624
                               19951121
                         Α2
                                         JP 1994-130773
                                                                  19940509 <--
PRAI JP 1994-130773
                               19940509 <--
    Antiaging cosmetics contain natural salt, chitosan (CM chitin) and
    polypeptides with addition of colorants, perfumes, and/or
     pharmaceutical natural products. A skin cosmetic contained natural salt
     comprising NaCl 85.0, Na2SO4 8.5, MgSO4 6.5, CaO and K2O ≤0.2 each,
     and harmful As, Cd, Cu, Pb, Zn and Br ≤ 0.01% in addition to other
     active ingredients and base materials.
ΙT
     9012-76-4, Chitosan 52519-63-8, Carboxymethylchitin
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
        (antiaging cosmetics containing natural salt, polypeptide and
        chitosan)
L115 ANSWER 19 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN
     1995:918401 HCAPLUS
DN
ΤI
     Novel drug delivery system by chitin derivative
ΑU
     Tokura, Seiichi; Nishi, Norio; Takahashi, Kiyohisa; Shirai, Akihiro;
     Uraki, Yasumitsu
     Graduate School of Environmental Earth Science, Hokkaido University,
CS
     Sapporo, 060, Japan
SO
     Macromolecular Symposia (1995), 99(Functional Polysaccharides),
     201-8
     CODEN: MSYMEC; ISSN: 1022-1360
PΒ
     Huethig & Wepf
DT
     Journal
LA
    Enalish
AΒ
    A porous chitin foam was regenerated from chitin dope in calcium chloride
     dihydrate saturated methanol. The porous chitin foam was shown to have
     cationic property, because chitin foam tended to adsorb anionic dyes
     through ionic binding and hydrophobic interaction. A pendant type of
     polymeric drug was prepared applying peptide spacer composed of
     phenylalanine at amino end and two step hydrolysis of polymeric drug were
     shown to release active drug at the final step using lysozyme and
     chymotrypsin in vitro.
IT
     1398-61-4, Chitin 52519-63-8, O-Carboxymethyl chitin
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (porous chitin foam as controlled-release drug delivery system)
L115 ANSWER 20 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN
AN
    1994:534814 HCAPLUS
DN
     121:134814
TΙ
     Preparation of cell-adhesive peptide bonded to polysaccharides
     Mori, Hideto; Komazawa, Hiroyuki; Saiki, Ikuo; Azuma, Ichiro
IN
PA
     Fuji Photo Film Co Ltd, Japan
     Jpn. Kokai Tokkyo Koho, 9 pp.
SO
     CODEN: JKXXAF
DT
     Patent
     Japanese
LA
FAN.CNT 1
                                         APPLICATION NO.
     PATENT NO.
                       KIND DATE
                                                                DATE
                                           _____
                               19940510 JP 1992-280292
     JP 06128289
                        A2
                                                                19921019 <--
PΤ
PRAI JP 1992-280292
                              19921019 <--
    MARPAT 121:134814
    Polysaccharides bonded to peptides X-Tyr-Ile-Gly-Ser-Arg-Y (X =
     absent, Glu, Asp; Y = NR1R2; R1, R2 = H, C1-4 alkyl) are prepared, which
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contain cell-adhesive core sequence of cell adhesive protein laminin. Typical polysaccharides are chondroitin sulfate, hyaluronic acid, and

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({\tt carboxymethyl}) \verb|chitin|. These \verb|peptide-polysaccharide| conjugates|
    retain various biol. activities of laminin, show high serum stability,
    more potent cell adhesiveness than the core sequence of laminin, and
    little side-effects, and are useful as cancer metastasis inhibitors.
    Thus, H-Tyr-Ile-Gly-Ser-Arg-NHCHMe2.2AcOH (I) was prepared by the solution
    method and condensed with carboxymethyl chitin by using
    1-ethyl-3-(3-dimethylaminopropyl)carbodiimide in 200 mM phosphate buffer
     (pH 7.4) to give I-carboxymethyl chitin conjugate containing 23 weight%
    peptide. In cancer metastasis assay, the latter
    glycopeptide reduced number of colonies of B16-BL6 melanoma cells
    formed in lungs of mice from 177±28 (control group) to 17±9.
TΤ
    52519-63-8DP, Carboxymethyl chitin, conjugate with laminin
    cell-adhesive core sequence-related peptide
    RL: SPN (Synthetic preparation); PREP (Preparation)
        (preparation of, as cell adhesion and cancer metastasis inhibitor)
L115 ANSWER 21 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN
    1994:525236 HCAPLUS
AN
DN
    121:125236
TI
    Water-soluble chitin or chitosan for treatment of arthritis
ΙN
    Nakagawa, Akira; Myata, Satoru; Shimozono, Juji; Soejima, Yoshiomi; Saida,
    Masaru
PΑ
    Hisamitsu Pharmaceutical Co, Japan
SO
    Jpn. Kokai Tokkyo Koho, 4 pp.
    CODEN: JKXXAF
DT
    Patent
LA
    Japanese
FAN.CNT 1
    PATENT NO.
                      KIND DATE
                                         APPLICATION NO.
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                                         _____
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                                                                  -----
                        A2 19940419 JP 1992-286967
    JP 06107551
                                                                19920930 <--
PRAI JP 1992-286967
                              19920930 <--
    Water-soluble chitin [e.g. carboxymethylchitin (I), dihydroxypropylchitin,
    hydroxyethylchitin, and carboxylchitin] or water-soluble chitosan (e.g.
    chitosan oligosaccharides, N-succinylchitosan, and hydroxypropylchitosan)
    are useful for treatment of arthritis. Administration of solution containing
    0.5% I to the knee joints showed greater analgesic effect in
    bradykinin-administered rats than that of a control containing Na hyaluronate.
    9012-76-4D, Chitosan, enzymic hydrolyzate 9056-32-0
    52519-63-8, Carboxymethylchitin 78809-92-4,
    N-Succinylchitosan 84069-44-3, Hydroxypropylchitosan
    84617-10-7, Dihydroxypropylchitin
    RL: BIOL (Biological study)
        (arthritis treatment with)
L115 ANSWER 22 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN
    1994:473202 HCAPLUS
ΑN
      Correction of: 1994:182560
DN
    121:73202
      Correction of: 120:182560
ΤI
    Synthesis of a MDP analog/chitin conjugate that stimulates cultured
    macrophages
ΑU
    Ohya, Yuichi; Murata, Junichi; Nishimoto, Takehiro; Ouchi, Tatsuro
    Fac. Eng., Kansai Univ., Suita, 564, Japan
CS
SO
     Journal of Bioactive and Compatible Polymers (1993), 8(4),
     351-64
    CODEN: JBCPEV; ISSN: 0883-9115
DT
    Journal
LA
    English
AΒ
    To provide a new synthetic biol. response modifier which exhibits a high
```

white - 10 / 810742 immunopotentiation activity and antitumor activity, a hybrid conjugate of chitin with immobilized D-glucose analog of muramyl dipeptide (MDP) (GADP) was synthesized. The stimulation activity of the conjugate against cultured macrophages was evaluated as an immunopotentiation activity in vitro by glucose consumption using PMA (phorbol-12-myristate-13-acetate)-differentiated HL-60 (human promyelocytic leukemia) cells and by superoxide anion (O2-) production from PMA-differentiated HL-60 cells. stimulation activity of the GADP/chitin conjugate against cultured macrophages was greater than that of GADP derivative, carboxymethyl-chitin and a mixture of these two. The stimulation activity of GADP against cultured macrophages was increased by conjugation with chitin. 52519-63-8DP, conjugates with MDP analogs RL: SPN (Synthetic preparation); PREP (Preparation) (preparation of, for tumor immunotherapy) L115 ANSWER 23 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN 1994:218328 HCAPLUS 120:218328 Design of D-glucose analog of MDP/polysaccharide conjugates and their immunological enhancement activities Ouchi, T.; Ohya, Y. Fac. Eng., Kansai Univ., Suita, 564, Japan New Funct. Mater. (1993), Volume B, 181-8. Editor(s): Tsuruta, Teiji. Publisher: Elsevier, Amsterdam, Neth.

- CODEN: 59NKAJ
- DT Conference
- English LA

ΤТ

AN

DN

TΙ

ΑU

CS

SO

- AB Hybrid type conjugates of or carboxymethylcurdlan carboxymethylchitin immobilizing D-glucose analog of N-acetylmuramyl-L-alanyl-D-isoglutamine were prepared and their immunol. enhancement activities were described.
- IT 52519-63-8, Carboxymethylchitin RL: RCT (Reactant); RACT (Reactant or reagent)

(coupling with glycopeptide)

- TΤ 52519-63-8DP, reaction products with muramoyldipeptide glucose analog
 - RL: SPN (Synthetic preparation); PREP (Preparation) (preparation and immunol. activity of)
- L115 ANSWER 24 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN
- 1993:633997 HCAPLUS AN
- DN 119:233997
- TISelective adsorption of peptides to carboxymethylated chitin
- ΑU Miura, Y.; Kaneda, Y.; Matsubara, N.; Nakano, H.; Tokura, S.
- CS Fac. Sci., Hokkaido Univ., Sapporo, 060, Japan
- SO Front. New Horiz. Amino Acid Res., Proc. Bienn. Int. Conf., 1st (1992), Meeting Date 1991, 385-9. Editor(s): Takai, Katsuji. Publisher: Elsevier, Amsterdam, Neth. CODEN: 59HEA5
- DT Conference
- LA English
- A 6-O-carboxymethyl-chitin (CM-chitin), one of the biodegradable derivs. AB from mucopolysaccharide, was studied as a carrier for sustained release of drugs by applying its several advantages for drug delivery system. The peptides containing neutral amino acid residue, especially phenylalanine (Phe), were found to be adsorbed specifically and stabilized by entrapping until CM-chitin was biodegraded to oligomers of small size.
- ΙT 52519-63-8, Carboxymethyl chitin RL: PEP (Physical, engineering or chemical process); PROC (Process) (peptides adsorption by, drug delivery in relation to)

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L115 ANSWER 25 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN
     1993:610434 HCAPLUS
AN
DN
     119:210434
TI
     Design of water-soluble CM-chitin/antitumor drug conjugate
ΑU
     Ouchi, Tatsuro; Inosaka, Keigo; Murata, Junichi; Nishimoto, Takehiro;
     Ohya, Yuichi
CS
     Fac. Eng., Kansai Univ., Suita, 564, Japan
SO
     Polymer Preprints (American Chemical Society, Division of Polymer
     Chemistry) (1992), 33(2), 537-8
     CODEN: ACPPAY; ISSN: 0032-3934
DT
     Journal
LA
     English
AB
     Conjugates of 5-fluorouracil with carboxymethyl chitin showed decreased
     side effects and conjugates of the glucose analog of
     muramylalanylisoglutamine with CM-chitin showed increased immunol.
     activity.
ΙT
     52519-63-8DP, conjugates with fluorouracil or muramyl
     dipeptide glucose analog
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); SPN (Synthetic preparation); BIOL (Biological
     study); PREP (Preparation)
        (preparation and biol. activity of)
L115 ANSWER 26 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN
ΑN
     1993:546447 HCAPLUS
DN
     119:146447
ΤI
     Stabilization of drug-linked peptides by 6-0-carboxymethyl
     Miura, Yoshiaki; Kaneda, Yoshihiro; Uraki, Yasumitsu; Tokura, Seiichi
ΑU
     Fac. Sci., Hokkaido Univ., Sapporo, 060, Japan
CS
     Adv. Chitin Chitosan, [Proc. Int. Conf.], 5th (1992), Meeting
SO
     Date 1991, 116-24. Editor(s): Brine, Charles J.; Sanford, Paul A.;
     Zikakis, John P. Publisher: Elsevier, London, UK.
     CODEN: 58YVAW
DT
     Conference
LA
     English
    While studying the specific adsorption of a model drug containing
AΒ
    phenylalanine to 6-0-carboxymethyl chitin (CM-chitin)-Ca2+ complex, it was
     observed that CM-chitin loses its adsorption capacity for the model drug, and
     releases the drug when it is degraded by lysozyme (one step release).
     When prodrugs were adsorbed into the CM-chitin-Ca2+ complex or were
     attached to CM-chitin through an enzyme susceptible spacer, the degradation of
     the CM-chitin backbone (first step) resulted in a sustained release of the
    prodrug. In the second step, the prodrug was converted to an active drug
     by a second enzymic hydrolysis (two step release). In the study of the
     pendant type of polymeric prodrug, it was also shown that the
     susceptibility for the second enzyme was enhanced with the decrease of the
    mol. weight of CM-chitin due to lysozymic hydrolysis.
ΙT
     52519-63-8 52519-63-8D, calcium complexes
     RL: BIOL (Biological study)
        (stabilization of peptide drugs by, prodrugs in relation to)
L115 ANSWER 27 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN
    1992:639902 HCAPLUS
DN
     117:239902
TI
     Ionically-cross-linked carboxyl-containing polysaccharides for
    post-operative adhesion prevention
ΙN
     Huang, W. James; Johns, Douglas B.; Kronenthal, Richard L.
PΑ
    Ethicon Inc., USA; Lifecore Biomedical, Inc.
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SO

Eur. Pat. Appl., 9 pp.

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CODEN: EPXXDW
DΤ
    Patent
LA
    English
FAN.CNT 2
    PATENT NO.
                   KIND DATE APPLICATION NO. DATE
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                                                            -----
    EP 507604
                      A2
                             19921007 EP 1992-302953
PΤ
                                                           19920403 <--
    EP 507604
                       A3
                             19931006
    EP 507604
                      В1
                             20050720
       R: AT, BE, CH, DE, ES, GB, IT, LI, LU, NL
    AU 9214015 A1
                             19921008 AU 1992-14015
                                                          19920402 <--
    AU 647905
                       B2
                             19940331
                   AA
                             19921006 CA 1992-2065111
    CA 2065111
                                                             19920403 <--
                      С
    CA 2065111
                             19991109
    AT 299706 E 20050815 AT 1992-302953 19920403 <-- EP 1593394 A2 20051109 EP 2005-76407 19920403 <--
       R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE
                A 19921201 BR 1992-1215 19920406 <--
A2 19930521 JP 1992-112371 19920406 <--
    BR 9201215
    JP 05124968
    JP 2702641
                       B2
                             19980121
PRAI US 1991-680955 A
EP 1992-302953 A3
                             19910405 <--
                            19920403 <--
AΒ
    Post-operative adhesion is reduced by topical application of a ionically
    cross-linked carboxyl-containing polysaccharide, such as CM-cellulose,
    carboxymethylchitin, hyaluronic acid, or their salts with alkali or
    alkaline-earth metals. The crosslinking agents are FeCl3, AlCl3, Al2(SO4)3 or
    Cr2(SO4)3. The composition may also contain inflammation inhibitors,
    growth factors or antibiotics.
ΙT
    52519-63-8D, Carboxymethylchitin, cross-linked, ionically
    RL: USES (Uses)
       (post-surgical adhesion prevention by)
L115 ANSWER 28 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN
    1990:406741 HCAPLUS
DN
ΤI
    Preparation of acidic polysaccharide esters as medicaments or
    biodegradable plastic materials, or for pharmaceutical vehicles and
    cosmetic preparations
ΙN
    Della Valle, Francesco; Romeo, Aurelio
    Fidia S.p.A., Italy
PΑ
SO
    PCT Int. Appl., 90 pp.
    CODEN: PIXXD2
DT
    Patent
LA
    Enalish
FAN.CNT 1
                  KIND DATE APPLICATION NO.
    PATENT NO.
                                                           DATE
    WO 8910940 A1 19891116
                      A1 19891116 WO 1989-EP520 19890512 <--
PΙ
       W: AU, DK, FI, HU, JP, KR
    EP 342557 A1 19891123
                                        EP 1989-108628
                                                              19890512 <--
                             19941123
                       В1
       R: AT, BE, CH, DE, ES, FR, GB, GR, IT, LI, LU, NL, SE
    AU 8935718 A1
                             19891129
                                        AU 1989-35718
                                                              19890512 <--
    AU 629551
                       B2
                             19921008
    HU 53127
                      A2
                             19900928
                                       HU 1989-3005
                                                              19890512 <--
    HU 208440
                   B 19931020
T2 19901129
B2 19991006
A 19920616
A2 19940921
                      В
                             19931028
    JP 02504164
                                      JP 1989-505459
                                                             19890512 <--
    JP 2958373
    US 5122598
                                        US 1989-350920
                                                              19890512 <--
    EP 615979
                                       EP 1994-107393
                                                             19890512 <--
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EP 615979
                         Α3
                               19941228
        R: AT, BE, CH, DE, ES, FR, GB, GR, IT, LI, LU, NL, SE
     ES 2063779
                        Т3
                               19950116
                                        ES 1989-108628
                                                                 19890512 <--
     CA 1336087
                        A1
                               19950627
                                           CA 1989-599556
                                                                 19890512 <--
                        A1
     IL 90273
                               19951127
                                           IL 1989-90273
                                                                19890512 <--
     DK 9000108
                        Α
                               19900312
                                           DK 1990-108
                                                                19900112 <--
     US 5466461
                        Α
                               19951114
                                           US 1992-862370
                                                                19920402 <--
PRAI IT 1988-47963
                       Α
                               19880513 <--
     EP 1989-108628
                        A3
                               19890512 <--
     US 1989-350920
                        A3
                               19890512 <--
     WO 1989-EP520
                         Α
                               19890512 <--
AΒ
     Total and partial esters of acidic polysaccharides chosen from
     (carboxymethyl)cellulose (I), -starch, and -methylchitin with aliphatic,
     araliph., or cycloaliph alcs. including pharmacol. active substances (e.g.
     alkaloids, anticonvulsants, and analgesics) and salts of such partial
     esters with (in)organic bases including therapeutically active amines (e.g.
    peptides, alkaloids, hormones, vitamins, and antivirals) are
     prepared They are useful as medicaments or biodegradable plastic materials
     for the preparation of sanitary and surgical articles (e.g. artificial skin in
     dermatol., surgical suture threads and sponges), as pharmaceutical
     vehicles (e.g. capsules for s.c. implant of medicament, microcapsules for
     s.c., i.m. or i.v. injection, and in other fields such as cosmetics, food
     or paper industry, adhesive products, etc. Thus, Bu4N+ salt of I (5.15 q)
     with a 0.75 substitution rate and medium viscosity was solubilized in DMSO
     at 25° with agitation and 1.71 g PhCH2Br was added and the solution
     stirred overnight at 30° to give, after precipitation and washing with
     EtOAc, 3.04g benzyl ester of I.
TΤ
    127565-90-6P
     RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
     (Reactant or reagent)
        (preparation and esterification of)
ΙT
     126041-92-7P 126041-93-8P 126041-94-9P
     126041-95-0P 126601-85-2P
     RL: SPN (Synthetic preparation); PREP (Preparation)
        (preparation of, as biodegradable plastic and pharmaceutical vehicle and for
        cosmetics)
TΤ
     52519-63-8DP, Carboxymethylchitin, esters
     RL: SPN (Synthetic preparation); PREP (Preparation)
        (preparation of, as biodegradable plastics and pharmaceutical vehicles and
        for cosmetics)
L115 ANSWER 29 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN
AN
    1990:406740 HCAPLUS
DN
     113:6740
ΤI
     Preparation of crosslinked carboxy polysaccharides as biodegradable
     plastic materials for cosmetics and pharmaceuticals
IN
     Della Valle, Francesco; Romeo, Aurelio
PA
     Fidia S.p.A., Italy
SO
     Eur. Pat. Appl., 37 pp.
     CODEN: EPXXDW
DT
     Patent
LA
    English
FAN.CNT 1
                                        APPLICATION NO.
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                               DATE
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    EP 341745
PΙ
                        A1
                               19891115
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        R: AT, BE, CH, DE, ES, FR, GB, GR, IT, LI, LU, NL, SE
     WO 8910941
                        A1
                              19891116 WO 1989-EP519
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W: AU, DK, FI, HU, JP, KR

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AU 8935747
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                                19891129
                                            AU 1989-35747
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    AU 631125
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    HU 53666
                          A2
                                19901128
                                            HU 1989-3636
                                                                    19890512 <--
    HU 210926
                          В
                                19950928
    JP 02504163
                          T2
                                19901129
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                                                                    19890512 <--
    JP 2941324
                          В2
                                19990825
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    EP 614914
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    DK 175386
                          В1
                                20040920
    FI 107050
                          В1
                                20010531
                                            FI 1990-188
                                                                    19900112 <--
    US 5676964
                          Α
                                19971014
                                            US 1995-465055
                                                                    19950605 <--
    GR 3034651
                          Т3
                                20010131
                                            GR 2000-402339
                                                                    20001023 <--
PRAI IT 1988-47964
                          Α
                                19880513
                                          <--
    EP 1989-108630
                          A3
                                19890512
                                          <--
    JP 1989-505458
                          A3
                                19890512 <--
    US 1989-350919
                          В1
                                19890512 <--
    WO 1989-EP519
                          Α
                                19890512 <--
    US 1993-70505
                          A1
                                19930601 <--
AB
     Inter- and/or intramol. esters of acid polysaccharides containing carboxy
     functions (e.g. auto-crosslinked polysaccharides), wherein (1) a first
    portion or all of the carboxy groups are esterified with hydroxy groups of
    the same mol. and/or of different mols. of the acid polysaccharide and/or
     (2) a second portion of the carboxy groups are esterified with a mono- or
    polyvalent alcs. including various drugs (e.g. alkaloids, anesthetic,
    analgesic, antiinflammatory, antiviral, antibacterial, etc.) and
    optionally salified with an alkali or alkaline earth metal, Mg, Al, or an
    amine including various drugs (e.g. alkaloids, peptides,
    antipsychotics, phenothiazine, vasoconstrictors, etc.), are prepared by
    treating an acidic polysaccharide (e.g., hyaluronic acid, alginic acid,
    CM-cellulose, carboxymethylchitin) with an activating agent (e.g.
     2-chloro-1-methylpyridinium iodide) and subjecting the resulting
    intermediate activated polysaccharide derivs. to heat or irradiation These
    auto-crosslinked polysaccharide acids are useful in the field of
    biodegradable plastic materials to manufacture sanitary and surgical articles
     (e.g. surgical suture thread, film for artificial skin, and sponges for
     the medication of wounds and lesions), for pharmaceutical vehicles for
     controlled-release of drugs (capsules for the s.c. implantation of
    medicaments or microcapsules for s.c., i.m., or i.v. injection), etc.
IT
    105156-94-3, Carboxymethylchitin sodium salt 127565-90-6
    RL: RCT (Reactant); RACT (Reactant or reagent)
```

(crosslinking of, by inter- and/or intramol. esterification) 52519-63-8DP, Carboxymethylchitin, cross-linked, sodium salt 52519-63-8DP, Carboxymethylchitin, cross-linked, sodium salt, Et

RL: SPN (Synthetic preparation); PREP (Preparation) (preparation of, for cosmetics, pharmaceutical vehicles, or medical goods)

L115 ANSWER 30 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN

1981:180713 HCAPLUS AN

DN

ester

ΤI Surgical lubricating powder for natural or synthetic rubber surgical

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elements
ΤN
     Casey, Donald James
PA
     American Cyanamid Co., USA
SO
     Brit., 9 pp.
     CODEN: BRXXAA
DΤ
     Patent
LA
     English
FAN.CNT 3
     PATENT NO.
                                   DATE APPLICATION NO.
                      KIND
                                                                      DATE
                                              -----
     _____
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                                  _____
PI GB 1583180 A 19810121 GB 1977-44643
US 4059097 A 19771122 US 1976-738502
US 4064564 A 19771227 US 1976-738200
US 4068757 A 19780117 US 1976-738501
BE 860423 A1 19780503 BE 1977-182300
PRAI US 1976-738200 A 19761103 <--
US 1976-738501 A 19761103 <--
US 1976-738502 A 19761103 <--
                                                                          -----
                                                                  19771026 <--
19761103 <--
19761103 <--
19761103 <--
19771103 <--
                                   19771227 US 1976-738200
                                   19780503 BE 1977-182300
AB
     A sterile surgical laminate package comprised a strippable laminate
     container containing a sterile rubber glove, on the surface of which was a
     lubricating powder consisting essentially of 1.5 g of an enzymically
     degradable form of poly(N-acetyl-D-glucosamine) (I) [27555-50-6]; the
     powder's particle size was 0.5\text{--}149~\mu and it would pass through a 200
     mesh screen. I was prepared by grinding com. chitin in a ball
     mill to a particle size of between 1 and 6 mm, followed by sequential
     treatment with 2N HCl, 90% HCO2H, and 10% NaOH. I could be used per se or
     converted into I membranes, poly[N-acetyl-6-0-(carboxymethyl
     )-D-glucosamine] [57216-53-2], poly[N-acetyl-6-0-(2'-
     hydroxyethyl)-D-glucosamine] [57216-54-3], or poly(N-acetyl-6-0-ethyl-D-
     glucosamine) [57216-56-5].
ΙT
     57216-53-2P
     RL: PREP (Preparation)
         (preparation of, as surgical glove lubricant)
L115 ANSWER 31 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN
ΑN
     1978:197685 HCAPLUS
DN
     88:197685
ΤI
     Chitin derived powder in sterile surgical element package
IN
     Casey, Donald James
PΑ
     American Cyanamid Co., USA
SO
     U.S., 8 pp.
     CODEN: USXXAM
     Patent
DT
LA
     English
FAN.CNT 3
                      KIND DATE APPLICATION NO.
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                                                                      DATÉ
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                                   -----
                                   19780117 US 1976-738501 19761103 <--
19790426 AU 1977-29651 19771013 <--
PΙ
     US 4068757
                          A
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     AU 7729651
     GB 1583180
                           A
                                   19810121
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                                                                        19771026 <--
                          A1
A
A
     DE 2748231
                                             DE 1977-2748231
                                   19780518
                                                                        19771027 <--
     SE 7712400
                                             SE 1977-12400
                                   19780503
                                                                        19771102 <--
     DK 7704873 A

JP 53058186 A2

NL 7712138 A

FR 2369826 A1

US 1976-738200 A
                                   19780504
                                              DK 1977-4873
                                                                        19771102 <--
                                               JP 1977-130946
                                   19780525
                                                                        19771102 <--
                                             NL 1977-12138
                                   19780508
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                                   19780602
                                               FR 1977-33071
                                                                        19771103 <--
PRAI US 1976-738200
                                   19761103 <--
     US 1976-738501 A
US 1976-738502 A
                                   19761103 <--
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19761103 <--

Natural or synthetic surgical goods are lubricated by a finely divided

AΒ

ΙT

ΙT

AN

DN

ΤI

ΙN

PΑ

SO

DT

LA

AΒ

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chitin-derived biodegradable powder of poly(N-acetyl-D-
      glucosamine) [27555-50-6], poly[N-acetyl-6-0-(carboxymethyl
      )-D-glucosamine [57216-53-2], poly[N-acetyl-6-0-ethyl-D-
      glucosamine [57216-56-5], or poly[N-acetyl-6-0-(2'-hydroxyethyl)-D-
      glucosamine [57216-54-3]. Lubricated gloves may be sterilized with no
      adverse effect on the disirable properties of the powder. The powder is
      readily absorbed by living tissue without deleterious tissue reaction.
      Thus, poly(N-acetyl-D-glucosamine) was obtained from powdered chitin
      by extraction with 2N HCl (decalcification), washing the material with water
      till neutral, and stirring it with 90% HCO2H overnight at room temperature
      mixture was centrifuged and water-washed residue was suspended in 10% NaOH
      and heated at 90-100^{\circ} for 2.5 h. The cake obtained after
      filtering, was washed with water until neutral and dried at 40°.
      1398-61-4D, hydrolyzates
      RL: BIOL (Biological study)
          (as surgical rubber lubricant)
      57216-53-2P
      RL: PREP (Preparation)
          (chitin derived surgical good lubricant, preparation of)
L115 ANSWER 32 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN
      1978:126373 HCAPLUS
      88:126373
      Minimizing tissue reaction during surgery with chitin
      Casey, Donald James
      American Cyanamid Co., USA
      U.S., 8 pp.
      CODEN: USXXAM
      Patent
      English
FAN.CNT 3
                        KIND DATE APPLICATION NO.
                                                                       DATE
PATENT NO.

PI US 4059097 A 19771122 US 1976-738502
AU 7729651 A1 19790426 AU 1977-29651
GB 1583180 A 19810121 GB 1977-44643
DE 2748231 A1 19780518 DE 1977-2748231
SE 7712400 A 19780503 SE 1977-12400
DK 7704873 A 19780504 DK 1977-4873
JP 53058186 A2 19780525 JP 1977-130946
NL 7712138 A 19780508 NL 1977-12138
FR 2369826 A1 19780602 FR 1977-33071
PRAI US 1976-738200 A 19761103 <--
US 1976-738501 A 19761103 <--
US 1976-738502 A 19761103 <--
      PATENT NO.
                                                 -----
                                     19771122 US 1976-738502 19761103 <--
19790426 AU 1977-29651 19771013 <--
19810121 GB 1977-44643 19771026 <--
                                                                             19771027 <--
                                                                             19771102 <--
                                                                             19771102 <--
                                                                             19771102 <--
                                                                             19771103 <--
                                                                             19771103 <--
      For diagram(s), see printed CA Issue.
      Surgical rubber gloves are lubricated by applying finely powdered
      biodegradable poly(N-acetyl-D-glucosamine) (I) [27555-50-6],
      poly[N-acetyl-6-0-(carboxymethyl
      )-D-glucosamine] [57216-53-2], poly[N-acetyl-6-
      O-(2'-hydroxyethyl)-D-glucosamine [57216-56-5], or poly[
      N-acetyl-acetyl-6-O-(ethyl)-D-glucosamine]
      [57216-54-3]. These powders were readily absorbed by living tissue
      without deleterious tissue reactions. The polymers were derived from
      chitin [1398-61-4]. Thus, finely ground com.
      chitin was decalcified by extracting with 2N HCl at 4° for 48 h.
      The material was collected by centrifugation and washed with water till
      neutral. The decalcified chitin was stirred at room temperature with
      HCO2H overnight. The mixture was centrifuged and the residue was washed
      with water. The washed chitin was suspended in 10% NaOH and was
```

heated at $90-100^{\circ}$ for 2.5 h. The solution was filtered, washed till neutral, and dried to give pure I. 57216-53-2 IT RL: PROC (Process) (as lubricant, for surgical rubber goods, preparation of) IT 1398-61-4 RL: BIOL (Biological study) (N-acetylglucosamine polymers derived from, for surgical rubber goods) L115 ANSWER 33 OF 33 HCAPLUS COPYRIGHT 2005 ACS on STN 1976:35314 HCAPLUS DN 84:35314 TΙ Enzymically decomposable bioerodible pharmaceutical carrier ΙN Capozza, Richard C. PA American Cyanamid Co., USA SO Ger. Offen., 24 pp. CODEN: GWXXBX DT Patent LA German FAN. CNT 1

PATENT NO. KIND DATE APPLICATION NO. DATE

OUS 3911098 A 19751007 US 1974-441695 19740211 <--
ZA 7500472 A 19760128 ZA 1975-472 19750122 <--
IL 46496 A1 19780831 IL 1975-46496 19750123 <--
AU 7577602 A1 19780729 AU 1975-77602 19750124 <--
GB 1499751 A 19780201 GB 1975-4193 19750130 <--
GB 1499751 A 19780201 GB 1975-4193 19750130 <--
NL 7501365 A 19750813 NL 1975-1365 19750205 <--
CA 1045975 A1 19790109 CA 1975-1365 19750205 <--
CA 1045975 A1 19790109 CA 1975-219603 19750207 <--
BE 825367 A1 19750811 BE 1975-153217 19750210 <--
SE 7501464 A 19750812 SE 1975-1464 19750210 <--
SE 7501464 A 19750812 SE 1975-1464 19750210 <--
FR 2260356 A1 19750905 FR 1975-4245 19750211 <--
DD 118801 C 19760320 DD 1975-184115 19750211 <--
ES 434618 A1 19770416 ES 1975-4245 19750211 <--
ES 434618 A1 19770416 ES 1975-860 19750211 <--
ES 434618 A1 19770421 <--
PRAI US 1974-441695 A 19740211 <--
AB An enzymically degradable form of poly (N-acetyl-D-glucosamine) (FAN.CNT 1 An enzymically degradable form of poly(N-acetyl-D-glucosamine) (chitin) [27555-50-6] served as a matrix for controlled release of drugs, especially in the eye. Degradable forms included also poly(N-acetyl-6-0carboxymethyl-D-glucosamine) [57216-53-2], poly[N-acetyl-6-0-(2-hydroxyethyl)-D-glucosamine] [57216-54-3], and poly(N-acetyl-6-O-ethyl-D-glucosamine) [57216-56-5], all of which were degraded by lysozyme [9001-63-2]. Preparation of these polymers from com. chitin was described. Films of the latter 3 polymers were prepared from aqueous solns.; suitable solvents for poly(N-acetyl-D-glucosamine) were hexafluoroacetone [684-16-2] sesquihydrate and hexafluoroisopropanol [920-66-1]. Thus, 50 mg pilocarpine nitrate [148-72-1] was added to a 5% aqueous solution of poly(N-acetyl-6-O-carboxymethyl-D-glucosamine) (0.95 g) and poured on a glass plate to form a 1.02 mm film which was dried and soaked in 10% alum solution for 5 hr. A 1 + 10 mm section of

IT 57216-53-2

RL: PRP (Properties)

caused pupil contraction lasting 6 hr.

(pharmaceutical controlled release from matrix of, in eye)

this film, placed on the eye surface of rabbits, was well tolerated and

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FILE LAST UPDATED: 14 NOV 2005 <20051114/UP>
MOST RECENT DERWENT UPDATE: 200573 <200573/DW>
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 http://thomsonderwent.com/support/userquides/ <<<
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 FOR FURTHER DETAILS: http://www.thomsonderwent.com/dwpifv <<<
- >>> THE CPI AND EPI MANUAL CODES HAVE BEEN REVISED FROM UPDATE 200501.
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- http://thomsonderwent.com/support/dwpiref/reftools/classification/code-revision/
 FOR DETAILS. <<<</pre>
- => d all abeq tech abex tot
- L150 ANSWER 1 OF 21 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN

AN 2005-655663 [67] WPIX

DNC C2005-198184

- TI Hydrogel composition useful for preventing the intrusion of micro-organisms into body cavities or body openings of mammals comprises a poly(N-vinyl lactam), a polysaccharide and water.
- DC A18 A25 A26 A96 B04 B05 C07 D21 D22
- IN BUONGIOVANNI, D; GRUENING, R; PERSCHBACHER, D J; QU, X; QU, X Y
- PA (HYDR-N) HYDROMER INC; (BUON-I) BUONGIOVANNI D; (GRUE-I) GRUENING R; (PERS-I) PERSCHBACHER D J; (QUXY-I) QU X Y

CYC 109

- PI US 2005191270 A1 20050901 (200567)* 11 A61K031-785 <--WO 2005086641 A2 20050922 (200567) EN A61K000-00 <--
 - RW: AT BE BG BW CH CY CZ DE DK EA EE ES FI FR GB GH GM GR HU IE IS IT KE LS LT LU MC MW MZ NA NL OA PL PT RO SD SE SI SK SL SZ TR TZ UG ZM ZW
 - W: AE AG AL AM AT AU AZ BA BB BG BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE EG ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NA NI NO NZ OM PG PH PL PT RO RU SC SD SE SG SK SL SM SY TJ TM TN TR TT TZ UA UG US UZ VC VN YU ZA ZM ZW
- ADT US 2005191270 A1 US 2004-788663 20040227; WO 2005086641 A2 WO 2005-US5323 20050218

PRAI US 2004-788663 20040227

IC ICM A61K000-00; A61K031-785 ICS A61K009-14

AB US2005191270 A UPAB: 20051019

NOVELTY - A hydrogel composition comprises a poly(N-vinyl lactam), a

polysaccharide and water (25 - 90), preferably 45 - 75, especially 55 - 65 weight%).

DETAILED DESCRIPTION - An INDEPENDENT CLAIM is included for a contraceptive hydrogel comprising a poly(N-vinyl lactam), a polysaccharide, water (25 - 55 weight%) and a spermicide. The weight ratio of the poly(N-vinyl) lactam to the polysaccharide is 75:1 - 1:5, 50:1 - 1:1 or 30:1 - 5:1.

ACTIVITY - Antimicrobial.

MECHANISM OF ACTION - Microbial growth inhibitor.

USE - For inhibiting the intrusion of micro-organisms into a body cavity e.g. natural body cavity (e.g. an ear canal, eye, nasal canal, mouth, genital opening, rectal opening, wrinkle or gland opening such as a teat canal of the milk gland of a dairy animal) or a cavity resulting from an injury (Claimed).

ADVANTAGE - The hydrogel compositions provide disinfecting/sanitizing activity without the need of antibiotics. Minimizing the use of antibiotics lowers the risk of antibiotic side effects, avoids long waiting periods after antibiotic applications and decreases the risk of developing antibiotic resistance in microorganisms. Moreover, as compared to current dry cow treatments which require complex processing steps, such as curing, and catalytic reactions, the hydrogel compositions are made by a simple mixing procedure. Also as compared to the current dry cow treatments, the hydrogel compositions are stable in a wide temperature range. The hydrogel compositions are biocompatible and lubricious. The hydrogel compositions have a consistency, which enable the hydrogel compositions to efficiently fill, and to remain in, body cavities/openings. Additionally, the consistency of these hydrogels allows for them to be squeezed out in total when needed or desired.

Dwq.0/0

FS CPI

MC

FA AB; DCN

CPI: A03-A01; A04-D05; A12-V01; A12-V03B; B01-B02; B03-A; B03-F; B03-H; B04-A08C; B04-A10; B04-B03A; B04-C02A1; B04-C02A2; B04-C02D; B04-C02E1; B04-C02E3; B04-C03; B04-N02; B05-A01A; B05-A01B; B05-A03A4; B05-A03A5; B05-A03B; B05-B01P; B05-B02C; B05-C07; B05-C08; B06-D03; B06-D09; B07-H; B09-D02; B10-A09B; B10-A12B; B10-A15; B10-A17; B10-B02J; B10-B04B; B10-C03; B10-C04D; B10-C04E; B10-D01; B10-D03; B10-E02; B10-E04B; B10-E04C; B10-E04D; B11-C02; B11-C04D; B12-M02; B12-M12C; B12-M12D; B14-A01; B14-A02; B14-A04; B14-C03; B14-L06; B14-P01A; C04-A08C; C04-C02A1; C04-C02A2; C04-C02D; C04-C02E1; C04-C02E3; C04-C03; C04-N02; C11-C02; C11-C04D; C12-M02; C12-M12C; C12-M12D; C14-A01; C14-A02; C14-A04; C14-C03; C14-L06; C14-P01A; D08-B; D09-A01; D09-A01C UPTX: 20051019

TECH

TECHNOLOGY FOCUS - INORGANIC CHEMISTRY - Preferred Composition: The hydrogel composition further comprises a radio-opaque additive. Preferred Components: The cross-linker is colloidal silica, colloidal alumina and/or colloidal titanium dioxide. The radio-opaque additive is barium sulfate, iodine contrast media or a tungsten particle. TECHNOLOGY FOCUS - ORGANIC CHEMISTRY - Preferred Composition: The composition additionally comprises a dye selected from a control dye, a food dye, a cosmetic dye, a FD and C dye or a D and C approved dye; a consistency modifying and/or performance modifying agent; and a radio-opaque additive.

Preferred Components: The vinyl monomer is selected from an acrylate, a hydroxyalkylacrylate, a methacrylate, an acrylic acid, a methacrylic acid and/or an acrylamide. The cross-linker is glutaraldehyde, genipin, aziridine derivative, carbimide derivative, epoxy, dialdehyde, paraformaldehyde and/or acrylamide. The consistency modifying and/or

performance modifying agent is methyl vinyl ether-co-maleic anhydride. The radio-opaque additive is iodine organic or bismuth organic.

TECHNOLOGY FOCUS - ORGANIC CHEMISTRY - Preferred Composition: The hydrogel composition further comprises a radio-opaque additive. Preferred Components: The cross-linker is colloidal silica, colloidal alumina and/or colloidal titanium dioxide. The radio-opaque additive is barium sulfate, iodine contrast media or a tungsten particle.

TECHNOLOGY FOCUS - PHARMACEUTICALS - Preferred Components: The hydrogel composition further comprises a therapeutic performance enhancing agent. Preferred Components: The therapeutic performance enhancing agent is an antimicrobial, antibacterial, antifungal, anti-candidiasis agent, disinfecting agent, biocide, bactericide, preservative, virucide, spermicide, germicide, sterilant, sanitizing ingredient, deodorizer, antiseptic, sporicide, a pharmaceutical, a veterinary preparation, an antibiotic, an anti-inflammatory agent, a plant or seed extract, a plant extract derivative, an herbal preparation and/or a humectant (preferably antimicrobial silver salt, silver zeolite, silver sulfadiazine, ethyl alcohol, isopropyl alcohol, benzyl alcohol, propionic acid, sorbic acid, salicylic acid, undecanoic acid, bleach, iodine, iodophor, potassium iodide, dodecyl benzene sulfonic acid, peroxide, bronopol, terbinafine, miconacole, econacole, clotrimazole, tolnaphthate, triclosan, trichlocarban, quaternary ammonium compound, benzalkonium halogenide, polyquatenium, polyquatenium derivative, formaldehyde releasing compound, hexetidin, chlorhexidine, chlorhexidine derivative, zinc pyrithione, zinc oxide, zinc propionate, paraben, phenoxyethanol, octoxynol-9, nonoxynol-9, ricinoleic acid, phenol mercuric acetate, sulfur lactic acid, essential oil of red thyme, allspice, cinnamon, savory, extract of rosemary, echinechea, nettle, fennel, juniper, ginseng, borage, gelsemium, hamamelis, poke root, arnica, aconite, apis, baptisia, thuja, aloe (barbadensis, vera, capensis), green tea, nasturtium, bryonia, eupatorium, and chamomile, acyclovir, idoxyumidine, ribavirin, vidarabine, rimantadine, aspirin, vitamin A and vitamin A derivative, vitamin E and vitamin E derivative, vitamin C and vitamin C derivative, betacarotin, betamethasone, dexamethasone, corticone and/or glycerin). Preferred Composition: The concentration of the therapeutic performance enhancing agent is 3, 7, 10, 15 or 20 wt.%. Water (15 - 75, 35 - 65 or 45 - 55 wt.%) is replaced by ethyl alcohol or isopropyl alcohol. TECHNOLOGY FOCUS - POLYMERS - Preferred Composition: The hydrogel composition further comprises a consistency modifying agent, a performance modifying agent and/or a cross-linker; and a radio-opaque additive. The poly(N-vinyl lactam) (0 - 5, 10, 20, 30, 40, 50, 60, 70, 80 or 90 wt.%) is replaced with the consistency and/or performance modifying copolymer. Preferred Components: The weight ratio of the upper boundary poly(N-vinyl) lactam to polysaccharide is 75:1; 50:1; 30:1; 20:1; 15:1; 13:1; 12:1; or 1:2. The weight ratio of the lower boundary poly(N-vinyl) lactam to polysaccharide is 1:10; 1:5; 1:3, 1:1; 5:1; 12:1; 13:1; 15:1; 20:1; 30:1;

lactam. The homopolymer is polyvinylpyrrolidone (PVP). The copolymer is a vinylpyrrolidone copolymer or an acrylamide copolymer. The terpolymer is a vinylpyrrolidone terpolymer, a vinylcaprolactam terpolymer or a dimethylaminoethyl methacrylate terpolymer.

The polysaccharide is chitin, deacetylated chitin, chitosan or its salt, chitosan sorbate, chitosan propionate, chitosan lactate,

terpolymer of N-vinyl lactam. The poly(N-vinyl lactam) is

or 50:1. The poly(N-vinyl lactam) is a homopolymer, a copolymer and/or a

N-vinylpyrrolidone, N-vinylbutyrolactam and/or N-vinylcaprolactam. The poly(N-vinyl lactam) is a vinyl monomer copolymerized with the N-vinyl

chitosan salicylate, chitosan pyrrolidone carboxylate, chitosan itaconate, chitosan niacinate, chitosan formate, chitosan acetate , chitosan gallate, chitosan glutamate, chitosan maleate, chitosan aspartate, chitosan glycolate, quaternary amine substituted chitosan salt, Ncarboxymethyl chitosan, ortho-carboxymethyl chitosan, N,-O-carboxymethyl chitosan, equivalent butyl chitosan derivative, cellulosic, alkylcellulose, nitrocellulose, hydroxypropylcellulose, starch or its derivative, methyl gluceth derivative, collagen, alginate, hyaluronic acid and/or heparin or its derivative. The consistency modifying and/or performance modifying agent is selected from polyvinyl alcohol, polyvinyl acetate, polyethylenoxide, poly(2-hydroxyethyl methacrylate), poly(ethylene-co-vinyl acetate), polyethylene glycol diacrylate, poly(N-isopropyl acrylamide), polyurethane, polyethylenimine, polypeptide, keratin, polyvinylpyrrolidone/polyethyleneimine, polyvinylpyrrolidone/polycarbamyl/polyglycol ester, polyvinylpyrrolidone/dimethylaminoethylmethacrylate/poly carbaml/polyglycol ester, polyvinylpyrrolidone/dimethiconylacrylate/polyca rbamyl/-polyglycol ester, or lecithin, or their copolymers and/or derivatives. The cross-linker is polyaminosilane, primary polyamine, polyaldehyde from acrolein reaction product and/or polyethylenimine. The radio-opaque additive is an iodine polymer. ABEX UPTX: 20051019 ADMINISTRATION - The hydrogel composition is applied by an injection device, infusion device, an applicator or plastic syringe (Claimed). No dosage given. EXAMPLE - Propylene glycol (1.4 g) and a 20% aqueous solution (3 g) of Pluronic F88 (RTM; a block copolymer of ethylene oxide and propylene oxide) were added to a 25% water solution (8.6 g) of Kollidon K90 (RTM; polyvinylpyrrolidone). To that solution, a 3% aqueous solution (5 q) of chitosan neutralized with pyrrolidone carboxylic acid were added. The mixture was stirred for a few minutes and transferred into plastic syringes for cavity applications. L150 ANSWER 2 OF 21 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN 2005-603255 [62] WPIX 1998-286610 [25]; 2004-118078 [12]; 2005-121248 [13]; 2005-372235 [38]; 2005-417021 [42]; 2005-424463 [43]; 2005-603254 [62] DNC C2005-181539 N2005-494765 Biological stent for the treatment of vascular atherosclerosis, includes body comprising crosslinked material having first degree of crosslink not less than the second degree of crosslink. A96 B05 D21 D22 P32 CHEN, M; SUNG, H; TU, H (CHEN-I) CHEN M; (SUNG-I) SUNG H; (TUHH-I) TU H CYC 1 US 2005163821 A1 20050728 (200562)* 56 A61F002-00 ADT US 2005163821 A1 CIP of US 2002-211656 20020802, CIP of US 2003-610391 20030630, CIP of US 2004-916170 20040811, CIP of US 2004-24101 20041228, US 2005-906239 20050210 FDT US 2005163821 A1 CIP of US 6624138 PRAI US 2005-906239 20050210; US 2002-211656 20020802; US 2003-610391 20030630; US 2004-916170 20040811; US 2004-24101 20041228 ICM A61F002-00

CR

DC ΙN

PΙ

IC

AΒ

US2005163821 A UPAB: 20050928

NOVELTY - A biological stent (41H) comprises a luminal surface portion

with a second degree of crosslink, an outer surface portion with a first degree of crosslink, and a body between the luminal and outer surface portions. The body comprises a crosslinked material having first degree of crosslink not less than the second degree of crosslink.

ACTIVITY - Vasotropic; Antiarteriosclerotic.

MECHANISM OF ACTION - None given.

USE - For the treatment of vascular atherosclerosis placing a biodegradable stent proximal to atherosclerosis, releasing bioactive agent; and treating vascular atherosclerosis distal to stent (claimed).

ADVANTAGE - The biological stent is biodegradable after serving its purpose, and has biocompatible breakdown products. It has also physical properties sufficient to perform its mechanical function. It has also sufficient longitudinal flexibility to facilitate insertion, and can deliver drugs locally to prevent restenosis.

DESCRIPTION OF DRAWING(S) - The figure shows an interlocking open-ring biodegradable stent of the invention.

Stent 41H

Member base 49

Ring elements 50A-B

Ends 53A-B

Dwg.18/21

FS CPI GMPI

FA AB; GI; DCN

MC CPI: A12-V03D; B04-C02B; B04-C02E; B04-C02E3; B04-C03; B04-H19; B04-N02; B05-A03B; B06-A02; B06-D18; B07-A04; B10-A20; B11-C04; B14-F02;

B14-F07; D08-B; D09-C01; D09-C04

TECH

UPTX: 20050928

TECHNOLOGY FOCUS - ORGANIC CHEMISTRY - Preferred Component: The crosslinked material is a biodegradable material from collagen, gelatin, elastin, chitosan, nitrogen-, oxygen-, carboxylmethyl chitosan (NOCC), low molecular weight (MW) chitosan, fibrin glue, biological sealant, chitosan -alginate complex, and/or chitosan-glycerol complex. It is crosslinked with a crosslinking material from genipin, its analog and/or derivatives, aglycon geniposidic acid, epoxy compounds, dialdehyde starch, glutaraldehyde, formaldehyde, dimethyl suberimidate, carbodiimides, succinimidyls, diisocyanates, acyl azide, and/or reuterin. The stent is sized and configured being a spiral or helical shape, a tubular mesh shape

or a non-tubular shape prior being loaded in a delivery apparatus. The

stent further comprises at least one bioactive agent.

TECHNOLOGY FOCUS - PHARMACEUTICALS - Preferred Component: The bioactive agent is analgesics or antipyretics, antiasthamatics, antibiotics, antidepressants, antidiabetics, antifungal agents, antihypertensive agents, anti-inflammatories, antineoplastics, antianxiety agents, immunosuppressive agents, antimigraine agents, sedatives or hypnotics, antipsychotic agents, antimanic agents, antiarrhythmics, antiarrhritic agents, antigout agents, anticoagulants, thrombolytic agents, antifibrinolytic agents, antiplatelet agents and antibacterial agents, antiviral agents, antimicrobials, and/or anti-infectives. It is conjugated to a targeting moiety from porphyrin or motexafin lutetium or a non-porphyrin drug facilitator.

TECHNOLOGY FOCUS - POLYMERS - Preferred Component: The biodegradable material can also be polylactic acid, polyglycolic acid, poly(D, L-lactide-co-glycolide), polycaprolactone, poly(amides), poly(ester amides), polyhydroxy acids, polyalkanoates, polyanhydrides, polyphosphazenes, polyetheresters, polyesteramides, polyesters, polyorthoesters, and/or their co-polymers.

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L150 ANSWER 3 OF 21 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN
ΑN
     2005-603254 [62]
                        WPTX
     1998-286610 [25]; 2004-118078 [12]; 2005-121248 [13]; 2005-372235 [38];
CR
     2005-417021 [42]; 2005-424463 [43]; 2005-580451 [59]; 2005-603255 [62]
DNN
    N2005-494764
                        DNC C2005-181538
TΙ
     Medical device, useful for delivering biological material to target
     tissues, comprises an apparatus having a surface, a bioactive agent and a
     biological material (comprising cross-linked bioactive agent) loaded onto
     portion of surface.
DC
     A96 B05 B07 D16 D22 P32
IN
     CHEN, M; LIANG, H; SUNG, H; TU, H
     (CHEN-I) CHEN M; (LIAN-I) LIANG H; (SUNG-I) SUNG H; (TUHH-I) TU H
PA
CYC
PΙ
     US 2005163818
                    A1 20050728 (200562)*
                                                23
                                                      A61F002-00
ADT
    US 2005163818 A1 Provisional US 1996-30701P 19961105, CIP of WO
     1997-US20113 19971104, CIP of US 2001-297808 20010927, CIP of US
     2002-211656 20020802, US 2003-610391 20030630
FDT
    US 2005163818 A1 CIP of US 6608040, CIP of US 6624138
PRAI US 1996-30701P
                          19961105; WO 1997-US20113
                                                         19971104:
     US 2001-297808
                          20010927; US 2002-211656
                                                         20020802;
     US 2003-610391
                          20030630
IC
     ICM A61F002-00
AB
     US2005163818 A UPAB: 20050928
     NOVELTY - Medical device (I) comprises:
          (a) an apparatus having a surface;
          (b) a bioactive agent; and
          (c) a biological material comprising the bioactive agent that is
     cross-linked with a cross-linking agent loaded onto at least a portion of
     the surface of (I).
          DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for:
          (1) a medical device being loaded with biological material being
     cross-linked with a cross-linking agent and a bioactive agent; and
          (2) a method for treating a target tissue of a patient comprising:
     cross-linking a biological material with a cross-linking agent; mixing the
     bioactive agent with the biological material; and delivering the
     biological material to the target tissue and releasing the bioactive agent
     for treating the target tissue.
          USE - (I) is useful for delivering the biological material to the
     target tissue and releasing the bioactive agent for treating the target
     tissue that comprises vulnerable plaque or atherosclerotic plaque (where
     the vulnerable plaque is the atherosclerotic plaque that is vulnerably
     prone to rupture) and the target tissue is tumor, cancer, brain tissue,
     vascular vessel or orthopedic tissue; preferably lymphatic vessel,
     gastrointestinal tract, hepatic duct, bile duct, pancreatic duct, urinary
     tract, ureter, urethra or reproductive tract (claimed).
          ADVANTAGE - The biological material is biodegradable or bioabsorbable
     for slow-release of the bioactive agent (claimed).
     Dwg.0/6
     CPI GMPI
FS
FΑ
     AB; DCN
     CPI: A12-V00V; B01-B02; B02-D; B04-A10; B04-C02B; B04-C02E;
MC.
          B04-C02E3; B04-E01; B04-F01; B04-H06; B04-N02; B04-N02A;
          B06-H; B07-A02B; B09-B; B10-A12C; B10-A15; B10-A20; B10-C04A;
          B10-C04B; B10-C04E; B10-D01; B11-C04; D09-C01
TECH
                    UPTX: 20050928
     TECHNOLOGY FOCUS - PHARMACEUTICALS - Preferred Components: The biological
     material is a solidifiable substrate, where (I) further comprises a step
     of solidifying the solidifiable substrate. The cross-linking agent is
     genipin, its analog and/or its derivatives; or formaldehyde,
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glutaraldehyde, dialdehyde starch, glyceraldehydes, cyanamide, diimides,

diisocyanates, dimethyl adipimidate, carbodiimide and/or epoxy compound. The apparatus is a stent or a non-stent implant (preferably annuloplasty rings, heart valve prostheses, venous valve bioprostheses, orthopedic implants, dental implants, ophthalmology implants, cardiovascular implants or cerebral implants; or a percutaneous device (a catheter, a wire, a cannula or an endoscopic instrument)). The biological material is: collagen, gelatin, elastin, chitosan, N, O, carboxylmethyl chitosan. The biological material is solidifiable from a phase (solution, paste, gel, suspension, colloid or plasma). The bioactive agent is: analgesics or antipyretics, antiasthmatics, antibiotics, antidepressants, antidiabetics, antifungal agents, antihypertensive agents, antiinflammatories, antineoplastics, antianxiety agents, immunosuppressive agents, antimigraine agents, sedatives or hypnotics, antipsychotic agents, antimanic agents, antiarhythmics, antiarthritic agents, antigout agents, anticoaqulants, thrombolytic agents, antifibrinolytic agents, antiplatelet agents and antibacterial agents, antiviral agents, antimicrobials or anti-infectives (preferably actinomycin D, paclitaxel, vincristin, methotrexate, angiopeptin, batimastat, halofuiginone, sirolimus, tacrolimus, everolimus, tranilast, dexamethasone, mycophenolic acid, lovastatin, thromboxane A2 synthetase inhibitors, eicosapentanoic acid, ciprostene, trapidil, angiotensin convening enzyme inhibitors, heparin, allicin, ginseng extract, flavone, ginkgo biloba extract, glycyrrhetinic acid, or proanthocyanides). The bioactive agent comprises biological cells (endothelial cells), genes or a growth factor (vascular endothelial growth factor, transforming growth factor-beta, insulin-like growth factor, platelet derived growth factor and/or fibroblast growth factor). The biological material is sized and configured as a medical device. Preferred Method: The method for treating a target tissue further comprises the step of chemically linking the bioactive agent with the biological material through a cross-linker before the solidifying step, where the bioactive agent comprises at least a cross-linkable functional group.

ABEX

UPTX: 20050928

ADMINISTRATION - Administration of (I) is via implantation.

EXAMPLE - Chitosan powder was dissolved in acetic acid at about pH 4. The deacetylation degree of the chitosan used was approximately 85%. The chitosan solution was adjusted to a pH of approximately 5.5 with sodium hydroxide. Drugs of interest were added into the chitosan solution. While loading the drug-containing chitosan onto the stent, the environment was adjusted to pH 7 with sodium hydroxide to solidify the chitosan onto the stent and it was further treated with a cross-linking agent (e.g. genipin) to enhance the biodurability and biocompatibility.

2005-458504 [46] ΑN WPIX DNC C2005-139340 TΙ Use of a topical composition comprising at least one compound of hydroxycarboxylic acids, N-acyl-aldosamines, Nacylamino acids and related compounds to enlarge mucocutaneous or cutaneous organs and sites. DC B05 D21 E19 ΙN VAN SCOTT, È J; YU, R J PΑ (VSCO-I) VAN SCOTT E J; (YURJ-I) YU R J

L150 ANSWER 4 OF 21 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN

CYC 108
PI WO 2005055947 A2 20050623 (200546)* EN 52 A61K000-00 <-RW: AT BE BG BW CH CY CZ DE DK EA EE ES FI FR GB GH GM GR HU IE IS IT

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KE LS LT LU MC MW MZ NA NL OA PL PT RO SD SE SI SK SL SZ TR TZ UG
            2M 2W
         W: AE AG AL AM AT AU AZ BA BB BG BR BW BY BZ CA CH CN CO CR CU CZ DE
            DK DM DZ EC EE EG ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG
            KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NA NI NO NZ
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     US 2005171194
                     A1 20050804 (200552)
                                                      A61K031-366
    WO 2005055947 A2 WO 2004-US41009 20041208; US 2005171194 A1 Provisional US
ADT
     2003-527307P 20031208, Provisional US 2004-570895P 20040514, US 2004-6822
     20041208
PRAI US 2004-570895P
                          20040514; US 2003-527307P
                                                         20031208;
     US 2004-6822
                          20041208
IC
     ICM A61K000-00; A61K031-366
     ICS A61K031-19
AB
    WO2005055947 A UPAB: 20050720
     NOVELTY - Enlarging mucocutaneous or cutaneous organs and sites comprises
     topically applying a composition (A) comprising at least one compound of
    hydroxycarboxylic acids, N-acyl-aldosamines, N-
     acylamino acids and related compounds for a period of time to the
    mucocutaneous or cutaneous organ or site.
          DETAILED DESCRIPTION - An INDEPENDENT CLAIM is also included for a
    method of preventing or ameliorating breast tumors comprising topically
     applying a composition, which comprises an antioxidant (citric acid,
     isocitric acid, malic acid, tartaric acid, pantolactone, isoascorbic acid,
     polyhydroxy acids, aldobionic acids or N-acetyl-cysteine) to the
     breast.
          ACTIVITY - Antiinflammatory; Analgesic; Anesthetic;
     Antibacterial; Virucide; Antifungal; Cytostatic; Dermatological;
     Endocrine-Gen.; Antiseborrheic; Antipruritic.
          MECHANISM OF ACTION - Histamine antagonist.
          USE - (A) is useful to: plump, pout, enhance or enlarge the lips and
     eyelids; plump, enhance or enlarge the breast; plump, enhance, enlarge
     and/or elongate the penis (claimed). The ability of (A) to enlarge or
     plump breast was tested in a female subject. The results showed that the
     breast had increased in plumpness and firmness after 3 months.
          ADVANTAGE - (A) has synergistic effect.
     Dwg.0/0
FS
    CPI
FA
    AB; DCN
MC
    CPI: B01-C05; B02-Z; B03-A; B05-A03A4; B07-A02; B07-D04A; B10-B02B;
          B10-C02; B10-C03; B10-C04B; B10-C04D; B10-E02; B12-M02B; B12-M07;
          B12-M12B; B14-A01; B14-A02; B14-A04; B14-C01; B14-C03; B14-C08;
          B14-D01; B14-H01E; B14-L09; B14-N17; B14-R05; B14-S08; B14-S09;
          D08-B09A; D08-B11; E07-A02B; E07-A02F; E07-D04A; E10-B02B; E10-C02A;
          E10-C02F; E10-C03; E10-C04B; E10-C04D4; E10-C04D5; E10-D01D;
          E10-E02F1; E35-C
TECH
                    UPTX: 20050720
     TECHNOLOGY FOCUS - ORGANIC CHEMISTRY - Preferred Components: The
    hydroxycarboxylic acid is alpha-hydroxyacids, beta-hydroxyacids,
     polyhydroxy acids and/or aldobionic acids. The hydroxycarboxylic
     acid is present as a free acid, salt, amide, ester and/or lactone. The
     alpha-hydroxyacid is alkyl alpha hydroxyacids, aralkyl alphahydroxyacids
     and/or polycarboxy alpha hydroxyacids. The polyhydroxy acid
     (PHAs) is an organic carboxylic acids having multiple hydroxyl
     groups in addition to the alphahydroxyl group, and where the polyhydroxy
     acid is present in the lactone form. The polyhydroxy acid is derived from
     carbohydrates and is aldonic acid, aldaric acid or alduronic acid. There
     are 9 hydroxycarboxylic acid compounds (B1) e.g. acid compounds
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of formulae R1R2C(OH)COOH (I), R1R2C(OH)COOH (II) and R1R2C(OH)COOH (III).

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The N-acyl-aldosamine (C1) of formula R1-(CHOH)m-CH(NHCOR2)-
(CHOH) n-R3. The N-acylamino acid (D1) is of formula
R1-CH(NHCOR2)-(CH2)n-COR3. The aldonic acid (E1) is of formula
R(CHOH)nCHOHCOOH. The aldaric acid (F1) is of formula HOOC(CHOH)nCHOHCOOH.
The alduronic acid (G1) is of formula HOOC(CHOH)nCHOHCHO. The aldobionic
acid (H1) is of formula H(CHOH)m(CHOR)(CHOH)nCOOH. In formula (I),
R1, R2 = H or alkyl (where the alkyl alpha hydroxyacid (AHA) can exist as
free acid, salt or partial salt with organic or inorganic alkali, amide,
ester, lactone, stereoisomers as D, L and DL or R, S and RS forms when R1
and R2 are not identical, and where the alkyl groups are non-aromatic
radicals such as methyl, ethyl, propyl, isopropyl, butyl, pentyl, octyl,
lauryl or stearyl).
In formula (II),
R1, R2 = H, aryl or aralkyl group (where the aralkyl AHAs are present as a
free acid, salt or partial salt with organic or inorganic alkali, amide,
ester, lactone, stereoisomers as D, L and DL or R, S and RS forms when R1
and R2 are not identical, and where the hydroxyl group is attached to a
non-aromatic alpha carbon atom).
In formula (III),
R1, R2 = H, COOH, CH2COOH or CHOHCOOH (where the polycarboxy AHA
are present as a free acid, salt or partial salt with organic or inorganic
alkali, amide, ester, lactone, stereoisomers as D, L and DL or R, S and RS
forms when R1 and R2 are not identical).
In formula (C1),
R1 = H, COOH, alkyl, alkoxyl, aralkyl and 1-19C aryl group;
R2 = alkyl, aralkyl or 1-19C group;
m, n = 0-19;
R3 = CHO, CONH2 or COOR4; and
R4 = H, an alkyl, aralkyl or 1-9C aryl group (where the hydrogen attached
to a carbon atom is optionally substituted by I, F, Cl, Br or alkyl,
alkoxyl, aralkyl or aryl group having 1 to 19 carbon atoms, and the N-
Acyl-aldosamine is present as a saturated or unsaturated,
stereoisomeric or non-stereoisomeric, straight or branched chain or cyclic
form).
In formula (D1),
Rl = H, an alkyl, aralkyl or 1-18C aryl;
R2 = alkyl, aralkyl or 1-18C aryl;
n = 0-5;
R3 = OH, NH2 OR3; and
R3 = alkyl, aralkyl or 1-9C aryl (in addition R1 optionally has OH, SH,
SCH3, COOH, NH2, CONH2, NHCONH2, NHC(=NH)NH2, imidazole, pyrrolidine or
other heterocyclic group, and the hydrogen attached to a carbon atom
optionally substituted by I, F, Cl, Br, OH or 1-9C alkoxyl).
In formula (E1),
R = H \text{ or alkyl group; and}
n = 1-6 (where the aldonic acid is present as a free acid, salt or
partial salt with organic or inorganic alkali, amide, ester, lactone, and
as a stereoisomer as D, L and DL or R, S and RS forms).
In formula (F1) and (G1),
n = 1-4 (where the aldaric acid and alduronic acid is present as a free
acid, salt or partial salt with organic or inorganic alkali, amide, ester,
lactone, or as a stereoisomer as D, L and DL or R, S and RS forms).
In formula (H1),
m, n = 0-7; and
R = monosaccharide (where the aldobionic acid is present as a free acid,
salt or partial salt with organic or inorganic alkali, amide, ester,
lactone, or as a stereoisomer as D, L and DL or R, S and RS forms).
The aldonic acid is 2,3-dihydroxypropanoic acid (glyceric acid);
2,3,4-trihydroxybutanoic acids (stereoisomers; erythronic acid and
erythronolactone, threonic acid and threonolactone); 2,3,4,5-
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tetrahydroxypentanoic acids (stereoisomers; ribonic acid and ribonolactone, arabinoic acid and arabinolactone, xylonic acid and xylonolactone, lyxonic acid and lyxonolactone); 2,3,4,5,6pentahydroxyhexanoic acids (stereoisomers; allonic acid and allonolactone, altronic acid and altronolactone, gluconic acid and gluconolactone, mannoic acid and mannolactone, gulonic acid and gulonolactone, idonic acid and idonolactone, galactonic acid and galactonolactone, talonic acid and talonolactone); and 2,3,4,5,6,7-hexahydroxyheptanoic acids (stereoisomers; alloheptonic acid and alloheptonolactone, altroheptonic acid and altroheptonolactone, glucoheptonic acid and glucoheptonolactone, mannoheptonic acid and mannoheptonolactone, guloheptonic acid and guloheptonolactone, idoheptonic acid and idoheptonolactone, galactoheptonic acid and galactoheptonolactone, taloheptonic acid and taloheptonolactone). The aldaric acid is 2,3-dihydroxybutane-1,4-dioic acids (stereoisomers; erythraric acid and threaric acid, also known as tartaric acid); 2,3,4-trihydroxypentane-1,5-dioic acids (stereoisomers; ribaric acid and ribarolactone, arabaric acid and arabarolactone, xylaric acid and xylarolactone, lyxaric acid and lyxarolactone); 2,3,4,5-tetrahydroxyhexane-1,6-dioic acids (stereoisomers; allaric acid and allarolactone, altraric acid and altrarolactone, glucaric acid and glucarolactone, mannaric acid and mannarolactone, gularic acid and gularolactone, idaric acid and idarolactone, galactaric acid and galactarolactone, talaric acid and talarolactone); 2,3,4,5,6pentahydroxyheptane-1,7-dioic acids (stereoisomers; alloheptaric acid and alloheptarolactone, altroheptaric acid and altroheptarolactone, glucoheptaric acid and glucoheptarolactone, mannoheptaric acid and mannoheptarolactone, guloheptaric acid and guloheptarolactone, idoheptaric acid and idoheptarolactone, galactoheptaric acid and galactoheptarolactone, taloheptaric acid and taloheptarolactone). The alduronic acid is erythruronic acid and threuronic acid; riburonic acid and riburonolactone; araburonic acid and araburonolactone; xyluronic acid and xyluronolactone; lyxuronic acid and lyxuronolactone; alluronic acid and alluronolactone; altruronic acid and altruronolactone; glucuronic acid and glucuronolactone; mannuronic acid and mannuronolactone; guluronic acid and guluronolactone; iduronic acid and iduronolactone; galacturonic acid and galacturonolactone; taluronic acid and taluronolactone; allohepturonic acid and allohepturonolactone; altrohepturonic acid and altrohepturonolactone; glucohepturonic acid and glucohepturonolactone; mannohepturonic acid and mannohepturonolactone; gulohepturonic acid and gulohepturonolactone; idohepturonic acid and idohepturonolactone; galactohepturonic acid and galactohepturonolactone; and talohepturonic acid and talohepturonolactone. The aldobionic acid is lactobionic acid and lactobionolactone from lactose, isolactobionic acid and isolactobionolactone from isolactose, maltobionic acid and maltobionolactone from maltose, isomaltobionic acid and isomaltobionolactone from isomaltose, cellobionic acid and cellobionolactone from cellobiose, gentiobionic acid and gentiobionolactone from gentiobiose, kojibionic acid and kojibionolactone from kojibiose, laminaribionic acid and laminaribionolactone from laminaribiose, melibionic acid and melibionolactone from melibiose, nigerobionic acid and nigerobionolactone from nigerose, rutinobionic acid and rutinobionolactone from rutinose, sophorobionic acid and sophorobionolactone from sophorose. The hydroxyacid is a hydroxyacid derivatives comprised of an ester form or an O-acetyl form of the hydroxyacid. The hydroxyacid derivative is glycolic acid methyl ester and ethyl ester, 0-acetyl-mandelic acid and 0-acetyl -benzilic acid. The hydroxyacid is a related hydroxycarboxylic acid selected from alpha ketoacids and miscellaneous hydroxyacids. The miscellaneous hydroxyacid is agaricic acid, aleuritic acid, citramalic acid, glucosaminic acid, galactosaminic acid, 2-keto-gulonic acid and

2-ketogulonolactone, mannosaminic acid, mevalonic acid and mevalonolactone, pantoic acid and pantolactone, quinic acid (1,3,4,5tetrahydroxycyclohexanecarboxylic acid), piscidic acid (4-hydroxybenzyltartaric acid), isoascorbic acid (D-erythro-hex-2-enonic acidr-lactone), 2-hexulosonic acids (isomers; arabino-2-hexulosonicacid, xylo-2-hexulosonic acid, ribo-2-hexulosonic acid, lyxo-2-hexulosonic acid), 5- hexulosonic acids (isomers; arabino-5-hexulosonic acid, xylo-5-hexulosonic acid, ribo-5-hexulosonic acid and/or lyxo-5-hexulosonic acid). The related compounds are related N-acetylamino acids such as N-acetyl-beta-alanine, N-acetyl -gamma-aminobutanoic acid, N-acetyl-beta-aminoisobutanoic acid, N-acetyl-citrulline, N-acetyl-dopa (N-acetyl -3,4-dihydroxyphenylalanine), N-acetyl-homocysteine, Nacetylhomoserine, N-acetyl-ornithine, N-acetyl -phenylglycine, N-acetyl-4-hydroxyphenylglycine or N, O-diacetyl-4-hydroxyphenylglycine. The Nacylamino acid is an N-propanoyllamino acid such as N-propanoyl-alanine, N-propanoyl-arginine, N-propanoyl-asparagine, N-propanoyl-aspartic acid, N-propanoyl-cysteine, N-propanoyl-glycine, N-propanoyl-glutamic acid, N-propanoyl-glutamine, N-propanoyl- histidine, N-propanoyl-isoleucine, N-propanoylleucine, N-propanoyl-lysine, N-propanoyl-methionine, N-propanoyl-phenylalanine, N-propanoyl-proline, N-propanoyl-serine, N-propanoyl-threonine, N-propanoyltryptophan, N-propanoyl-tyrosine or N-propanoyl-valine. The related compounds are related N-propanoylamino acids such as N-propanoyl-beta-alanine, N-propanoyl-gamma-aminobutanoic acid, N-propanoyl-beta-aminoisobutanoic acid, N-propanoyl-citrulline, N-propanoyl-dopa (N-propanoyl-3,4dihydroxyphenylalanine), N-propanoyl-homocysteine, N-propanoyl-homoserine, N-propanoyl-ornithine, N-propanoyl-phenylglycine, N-propanoyl-4hydroxyphenylglycine or N,O-dipropanoyl-4-hydroxyphenylglycine. The aldonic acid is about 0.01-99.9 (preferably 1-25) wt.% of (A). The effective period of time is for at least two weeks (preferably at least six months). (A) further comprises a cosmetic, pharmaceutical or other topical agent. The cosmetic, pharmaceutical or other topical agent is of agents that improve or eradicate age spots, keratoses and wrinkles; local analgesics and anesthetics; antiacne agents; antibacterials; antiyeast agents; antifungal agents; antiviral agents; antidermatitis agents; antihistamine agents; antipruritic agents; antiinflammatory agents; antipsoriatic agents; antiseborrheic agents; antiaging and antiwrinkle agents; sunblock and sunscreen agents; skin lightening agents; depigmenting agents; vitamins; corticosteroids; tanning agents; humectants; estrogens; androgens; hormones or retinoids. The cosmetic, pharmaceutical or other topical agent is aclovate, acyclovir, acetylsalicylic acid, adapalene, albuterol, aluminum acetate, aluminum chloride, aluminum hydroxide, aluminum chlorohydroxide, amantadine, aminacrine, aminobenzoic acid (PABA), aminocaproic acid, aminosalicylic acid, amitriptyline, anthralin, ascorbic acid, ascoryl palimate, atropine, azelaic acid, bacitracin, bemegride, beclomethasone dipropionate, benzophenone, benzoyl peroxide, betamethasone dipropionate, betamethasone valerate, brompheniramine, bupivacaine, butoconazole, calcipotriene, camphor, capsaicin, carbamide peroxide, chitosan, chlorhexidine, chloroxylenol, chlorpheniramine, ciclopirox, clemastine, clindamycin, clioquinol, clobetasol propionate, clotrimazole, coal tar, cromolyn, crotamiton, cycloserine, dehydroepiandrosterone, desoximetasone, dexamethasone, diphenhydramine, doxypin, doxylamine, dyclonine, econazole, erythromycin, estradiol, estrone, ethinyl estradiol, fluocinonide, fluocinolone acetonide, 5-fluorouracil, griseofulvin, guaifenesin, haloprogin, hexylresorcinol, homosalate, hydrocortisone, hydrocortisone 21 -acetate, hydrocortisone 17-valerate,

hydrocortisone 17-butyrate, hydrogen peroxide, hydroquinone, hydroqumone monoether, hydroxyzine, ibuprofen, ichthammol, imiquimod, indomethacin, ketoconazole, ketoprofen, kojic acid, lidocaine, meclizine, meclocycline, menthol, mepivacaine, methyl nicotinate, metronidazole, miconazole, minocycline, minoxidil, monobenzone, mupirocin, naftifine, naproxen, neomycin, nystatin, octyl methoxycinnamate, octyl salicylate, oxybenzone, oxiconazole, oxymetazoline, padimate O, permethrin, pheniramine, phenol, phenylephrine, phenylpropanolamine, piperonyl butoxide, podophyllin, podofilox, povidone iodine, pramoxine, prilocaine, procaine, promethazine propionate, propranolol, pseudoephedrine, pyrethrin, pyrilamine, resorcinol, retinal, 13-cis retinoic acid, retinoic acid, retinol, retinyl acetate, retinyl palmitate, selenium sulfide, shale tar, sulconazole, sulfur, sulfadiazine, tazarotene, testosterone, terbinafine, terconazole, tetracaine, tetracycline, tetrahydrozoline, thymol, tioconazole, tolnaftate, triamcinolone diacetate, triamcinolone acetonide, triamcinolone hexacetonide, triclosan, triprolidine, undecylenic acid, urea, vitamin E acetate, wood tar or zinc pyrithione. ABEX UPTX: 20050720 SPECIFIC COMPOUNDS - The use of 20 compounds is specifically claimed as (D1) e.g. N-acetyl-alanine, N-acetylarginine, Nacetyl-asparagine, N-acetyl-aspartic acid, Nacetyl-cysteine, N-acetylglycine, N-acetyl -glutamic acid, N-acetyl-glutamine, N-acetylhistidine and N-acetylisoleucine. The use of 108 compounds is specifically claimed as (C1) e.g. N-acetyl-glycerosamine, Nacetyl-erythrosamine, N-acetyl-threosamine, Nacetyl-ribosamine, N-Acetylarabinosamine, N-Acetyl-xylosamine, N-Acetyl-lyxosamine, N-Acetyl -allosamine, N-Acetyl-altrosamine, N-Acetyl -glucosamine, N-Acetyl-mannosamine, N-Acetylgulosamine , N-Acetyl-idosamine, N-Acetyl-galactosamine and N-Acetyl-talosamine. The use of 32 compounds is specifically claimed as (B1) e.g. 2-hydroxyethanoic acid (glycolic acid); 2-hydroxypropanoic acid (lactic acid); 2-methyl-2-hydroxypropanoic acid (methyllactic acid); 2-hydroxybutanoic acid; 2-hydroxypentanoic acid; 2,3-dihydroxybutane-1,4dioic acid (tartaric acid); 2-ketoethanoic acid (glyoxylic acid); 2-ketopropanoic acid (pyruvic acid) and 2-phenyl-2-ketoethanoic acid (benzoylformic acid). ADMINISTRATION - Administration of (A) is topical (claimed). L150 ANSWER 5 OF 21 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN 2005-384670 [39] WPIX DNC C2005-119027 Composition useful e.g., for treating a dental dry socket, and in wound dressings, comprises a hydrogel of a non-acidic poly(n-vinyllactam), a water-soluble multifunctional amine-containing polymer and/or a chitosan derivative. A18 A28 A96 B07 D21 HORNG, L L (HORN-I) HORNG L L 108 US 2005112151 A1 20050526 (200539) * 10 A61K007-06 WO 2005055924 A2 20050623 (200541) EN A61K000-00 <--RW: AT BE BG BW CH CY CZ DE DK EA EE ES FI FR GB GH GM GR HU IE IT KE LS LU MC MW MZ NA NL OA PL PT RO SD SE SI SK SL SZ TR TZ UG ZM ZW W: AE AG AL AM AT AU AZ BA BB BG BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE EG ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NA NI NO NZ

ΑN

ΤI

DC

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PA

ΡI

CYC

OM PG PH PL PT RO RU SC SD SE SG SK SL SY TJ TM TN TR TT TZ UA UG US UZ VC VN YU ZA ZM ZW

ADT US 2005112151 A1 US 2003-707102 20031120; WO 2005055924 A2 WO 2004-US28292 20040831

PRAI US 2003-707102 20031120

- IC ICM A61K000-00; A61K007-06
 - ICS A61K007-00; A61K007-11
- AB US2005112151 A UPAB: 20050621

NOVELTY - A composition (A) comprises a hydrogel formed by a mixture of two or more of (1) a non-acidic poly(n-vinyl lactam) with a k value of at least 30; (2) a water soluble multifunctional amine-containing polymer or its mixtures; and (3) a **chitosan** derivative or its mixtures.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for:

- (1) a composition (A) comprising (1) with a K value of at least 30 and (2) in a weight ratio of about 8.75-1;
- (2) a composition (B) comprising a hydrogel formed by a mixture of
 (1) and (2);
- (3) a composition (C) formed by combining an aqueous solution (a) comprising non-acidic polyvinylpyrrolidone, lidocaine hydrogen chloride and glutaric dialdehyde in a weight ratio, respectively, of about 69.7 to 12 to 1 with an aqueous solution (b) comprising polyethyleneimine, glycerin and polyvinylpyrrolidone/dimethylaminoethyl-methacrylate copolymer in a weight ratio, respectively, of about 1.87 to 1.25 to 1, where the total weight of aqueous solution of each of (a) and (b) are in the range of about 0.9 to 1 respectively;
- (4) a composition formed by combining an aqueous solution (c) comprising non-acidic polyvinylpyrrolidone, polyethylene glycol, benzocaine and glutaric dialdehyde in a weight ratio respectively of about 56 to 34 to 14 to 1 with an aqueous solution (d) comprising polyethylene glycol, glycerin, benzocaine, polyvinyl pyrrolidone/dimethylaminoethylmethacrylate copolymer and polyethyleneimine in the weight ratio respectively of about 5.5 to 2.5 to 2.5 to 2 to 1, where the total weight of each aqueous solution (c) and (d) is in the range of about 1 to 1;
- (5) a composition (D) comprising a hydrogel formed by a mixture of
 (1) with a K value of at least 30 and/or (3);
- (6) a composition (E) comprising a hydrogel formed by the mixture of
 (2) and/or (3);
- (7) a dental anesthetic application (F) comprising (B) further including an anesthetic (lidocaine, benzocaine and Eugenol (about 1-30 weight%), moisturizers and plasticizers (about 0-50 weight%) and preservatives (about 0-4 weight%) and where the weight ratio of (1) to (2) is about 80/1 to 2/1;
 - (8) a cosmetic face mask comprising (A);
- (9) a kit for a cosmetic gel (A) and a separate portion containing cosmetic agents (hydrating agents, fragrances and skin nutrients) with instructions to the order of addition to and the amount of water in which to form a hydrogel, application and removal directions; and
- (10) a hydrogel in sheet or roll form comprising (A) and further including a releasable backing sheet.
- USE The compositions (A-E) are useful to treat a dental dry socket (claimed). The compositions (A-E) are useful as wound dressings, burn dressings, drug delivery systems, cosmetic masks, conductive electrodes, prostheses and wraps. The hydrogel are useful as carriers for wide range of pharmaceutically acceptable and releasable biologically active agents having curative or therapeutic value for human or non-human animals.

ADVANTAGE - The compositions supply the patients with a less painful alternative to the standard gauze packing treatment. The ring opening of (1) of the compositions is omitted. The compositions are stable and economical. Dwg.0/0

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FS
     CPI
FA
     AB; DCN
MC
     CPI: A04-D05; A12-S; A12-V02B; A12-V03A; A12-V04C; B04-C03; B05-A03B;
          B06-D09; B06-E05; B07-A02A; B07-D09; B10-A05; B10-B03B; B10-C03;
          B10-E02; B10-E04A; B14-A01; B14-A02; B14-A04; B14-C01; B14-C03;
          B14-C08; B14-F01; B14-F02; B14-F02C; B14-H01; B14-J02A; B14-J05D;
          B14-N06; B14-R01; D08-B09A; D09-C04B; D09-C06
TECH
                    UPTX: 20050621
     TECHNOLOGY FOCUS - PHARMACEUTICALS - Preparation: (A) is prepared by
     reaction of (a) an aqueous solution comprising non-acidic
     polyvinylpyrrolidone and glycerin in a weight ratio of about 1.5 to 1 with
     (b) an aqueous solution comprising carboxymethyl
     chitosan, glycerin and polyethyleneimine in a weight ratio of
     about 6.6 to 156 to 1 respectively, where the total weight of aqueous
     solutions (a) and (b) are in the range of about 1 to 1.
     Preferred Components: (1) is homopolymers, coplymers and terpolymers of
     N-vinyl lactam. The copolymers and terpolymers of (1) are N-vinyl lactam
     monomer copolymerized with monomers containing a vinyl functional group.
     The vinyl containing monomers are acrylates, hydroxyalkylacrylates,
     methacrylates, acrylic acid, methacrylic acid, acrylamindes,
     vinylpyrrolidone, vinylcaprolactam or dimethylaminoethyl methacrylate
     terpolymers. The homopolymer of (1) is vinylpyrrolidone. (2) is
     polyethyleneimine, amine terminated polyethylene oxide polymers, amine
     terminated polyethylene/polypropylene oxide polymers, polymers and
     copolymers of dimethyl amino ethyl methacrylate or vinyl pyrrolidone. (3)
     is a biocompatible salt such as a chitosan reacted with a
     reactant (pyrrolidone carboxylic acid, glutamic acid or an
     acetate); preferably N, O-carboxymethyl
     chitosan or N, O-carboxybutyl
     chitosan. (A) further includes biologically active and
     pharmaceutically acceptable substances having curative or therapeutic
     value; an electrolyte (sodium chloride, potassium chloride or magnesium
     acetate), where the hydrogel is rendered electrically conductive;
     a skin-hydrating agent (water, sodium pyrrolidone carboxylate,
     lactic acid, hyaluronic acid or hydrolyzed collagen); enhancing agents
     (wetting agents, moisturizers, plasticizers, surfactants or dispersing
     agents; preferably glycerin, propylene glycol or polyethylene glycol); an
     electrolytic salt as an anti-osmotic agent (an alkali metal chloride or
     sodium bicarbonate); additives (polymer lattices, fillers, surfactants,
     pigments, dyes or fragrances). The biologically active materials are
     hypnotics, sedatives, tranquilizers, anti-convulsants, muscle relaxants,
     analgesics, antipyretic agents, anti-inflammatory
     agents, local anesthetics, antispasmodics, anti-ulcer agents, anti-virals,
     anti-bacterials, anti-fungals, sympathomimetic agents, cardiovascular
     agents or antitumor agents; preferably nitroglycerine, scopolamine,
     pilocarpine, ergotamine tartrate, phenylpropanolamine, theophylline,
     antimicrobials tetracycline, neomycin, oxytetracycline, triclosan, sodium
     cefazolin, silver sulfadiazine, methylsalicylate, salicylic acid,
     nicotinates, methyl nicotinate, chlorhexidine gluconate, menthol,
     capsicum, lidocaine or benzocaine. The weight ratio of poyvinylpyrrolidone
     to (2) is about 2/1 to 80/1. In the preparation of (C), the weight ratio
     of the aqueous solution comprising non-acidic polyvinylpyrrolidone,
     lidocaine hydrogen chloride and glutaric dialdehyde respectively, is about
     68.2 to 16 to 1 and the weight ratio of aqueous solution comprising
     polyethyleneimine, glycerin and polyvinyl pyrrolidone/dimethylaminoethyl-
     methacrylate copolymer, respectively, is about 1.85 to 1.25 to 1, where
     the total weight of the aqueous solution in each of (a) and (b) are in the
     range of about 1 to 1 respectively. The weight ratio of the poly(N-vinyl
     lactam) to chitosan derivative is in the range of from about 2/1
```

to about 100/1. The composition (D) comprises about 1-30 wt.% of

pharmaceutically acceptable local anesthetic, about 0-50 wt.% of a moisturizer, about 0-50 wt.% of a plasticizer, about 0-4 wt.% of a preservative, were the weight ratio of (1) to (3) is about 80/1 to about 2/1. In (D), the poly(N-vinyl lactam) is polyvinylpyrrolidone and the weight ratio of polyvinylpyrrolidone to (3) is about 17.5/1. In (E), the weight ratio of (2) and (3) is in the range of 50/1 to 1/50. In (E), (2)is polyethyleneimine and (3) is carboxymethyl chitosan and the weight ratio of the amine to the chitosan derivative is about 1.2-1 (preferably 0.3 to 1 or 0.15-1). (E) further includes about 13 (preferably 47) wt.% of glycerin. In (F), the eugenol is about 2-20 wt.%, the moisturizers and plasticizers is about 5-25 wt.% and the preservatives is about 0.01-2 wt.% and where the weight ratio of (1) to (2) is about 30/1 to 5/1. The releasable backing layer provides protection of the hydrogel from gases, liquid, air and the selection of area to be treated. Preferred Method: In the treatment of dry socket, (A) is applied to the socket as a layer. UPTX: 20050621 EXAMPLE - Nonacidic K60 PVP (79 g, 40 weight%) at pH 7 and glycerin (21 g) in aqueous solution was mixed with carboxymethyl chitosan (2 g), aqueous solution of glycerin (47 g), 50 weight% polyethyleneimine aqueous solution (0.6 g) and water (51 g) and a gel was formed in less than 5 minutes.

aqueous solution was mixed with carboxymethyl chitosan

(2 g), aqueous solution of glycerin (47 g), 50 weight% polyethyleneimine aqueous solution (0.6 g) and water (51 g) and a gel was formed in less than 5 minutes.

L150 ANSWER 6 OF 21 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN

AN 2005-372235 [38] WPIX

CR 1998-286610 [25]; 2004-118078 [12]; 2005-121248 [13]; 2005-417021 [42]; 2005-424463 [43]; 2005-580451 [59]; 2005-603254 [62]; 2005-603255 [62]

DNN N2005-301078 DNC C2005-115289

TI Crosslinked biodegradable stent/implant useful for treating target tissue e.g. vulnerable plaque, comprises at least one layer/zone of biological material crosslinked with crosslinking agent contains at least one bioactive agent.

DC A96 B07 D22 P32

IN CHEN, M; SUNG, H; TU, H; TU, P Y

PA (GPME-N) GP MEDICAL INC

CYC 108

ABEX

PI WO 2005046519 A1 20050526 (200538)* EN 62 A61F002-02

RW: AT BE BG BW CH CY CZ DE DK EA EE ES FI FR GB GH GM GR HU IE IS IT KE LS LU MC MW MZ NA NL OA PL PT RO SD SE SI SK SL SZ TR TZ UG ZM ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE EG ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NA NI NO NZ OM PG PH PL PT RO RU SC SD SE SG SK SL SY TJ TM TN TR TT TZ UA UG US UZ VC VN YU ZA ZM ZW

ADT WO 2005046519 A1 WO 2004-US37217 20041105

PRAI US 2004-585775P 20040706; US 2003-518050P 20031107; US 2004-547935P 20040226; US 2004-565438P 20040426; US 2004-574501P 20040526; US 2004-610391 20040630

IC ICM A61F002-02

AB W02005046519 A UPAB: 20050928

NOVELTY - A crosslinked biodegradable stent/implant (21) comprises at least one layer or zone (22A) of biological material (24). The biological material contains at least one bioactive agent and is crosslinked with crosslinking agent.

ACTIVITY - Antiarteriosclerotic. No suitable biodata given. MECHANISM OF ACTION - None given.

USE - For treating a target tissue such as atherosclerosis plaque or vulnerable plaque (claimed).

ADVANTAGE - Crosslinking of a drug-containing biological material

with genipin enables the resulting material with less antigenicity or immunogenicity. The implant/stent exhibits many of desired characteristics important for optimal therapeutic function. The implant/stent provides controlled and sustained release of drug over an extended period of time. DESCRIPTION OF DRAWING(S) - The figure shows biodegradable zone. Stent 21 First zones 22A, 22B Second zone 23 First biodegradable material 24 Portion of continuous circumference 25 Second biodegradable material 26 Dwg.9/18 CPI GMPI FS FΑ AB; GI; DCN MC CPI: A12-V02; B01-B02; B02-E; B02-S; B02-T; B04-B01B; B04-C02B; **B04-C02E3**; B04-H06; B04-H19; B04-N02; B05-B01G; B05-B01P; B06-H; B07-H; B09-B; B10-A04; B10-A17; B10-A20; B10-C03; B10-C04A; B10-D01; B11-C04A; B12-M10A4; B14-A01; B14-A02; B14-A04; B14-C01; B14-C02; B14-C03; B14-C09; B14-D10; B14-F01A; B14-F02B; B14-F02C; B14-F04; B14-F07; B14-F08; B14-H01; B14-J01B; B14-K01A; B14-S04; D09-C04

TECH

UPTX: 20050616

TECHNOLOGY FOCUS - BIOLOGY - Preferred Biological Material: The biological material is selected from collagen, gelatin, elastin, chitosan, N,O-carboxymethyl chitosan (

NOCC), fibrin glue, biological sealant and/or chitosan -alginate complex. The bioactive agent is ApoA-I Milano, recombinant ApoA-I Milano/phospholipid complexes, biological cells or epithelial progenitor cells or a growth factor. The growth factor is selected from vascular endothelial growth factor, transforming growth factor-beta, insulin-like growth factor, platelet derived growth factor, and/or fibroblast growth factor.

TECHNOLOGY FOCUS - PHARMACEUTICALS - Preferred Agent: The bioactive agent is selected from analgesics/antipyretics (e.g. aspirin), antiasthamatics, antibiotics, antidepressants, antidiabetics, antifungal agents, antihypertensive agents, anti-inflammatories,

antineoplastics (e.g. actinomycin D), antianxiety agents, immunosuppressive agents, antimigraine agents, sedatives/hypnotics, antipsychotic agents, antimanic agents, antiarrhythmics, antiarthritic agents, antigout agents, anticoagulants, thrombolytic agents, antifibrinolytic agents, antiplatelet agents and antibacterial agents, antiviral agents, antimorobials, and/or anti-infective or paclitaxel, methotrexate, angiopeptin, batimastat, halofuginone, sirolimus, tacrolimus, everolimus, ABT-578, tranilast, dexamethasone, mycophentanoic acid, lovastatin, thrombozane A2 synthetase inhibitors, eicosapentanoic acid, ciprostene, trapidil, angiotensin convening enzyme inhibitors, heparin, allicin, ginseng extract, ginsenoside Rgl, flavone, ginkgo biloba extract, glycyrrhetinic acid, lipostabil and/or proanthocyanides.

TECHNOLOGY FOCUS - ORGANIC CHEMISTRY - Preferred Crosslinking Agent: The biological material is crosslinked with crosslinking agent or reversible crosslinking agent (preferably poly(amides) or poly(ester amides)). The crosslinking agent is selected from aglycon geniposidic acid, epoxy compounds, dialdehyde starch, glutaraldehyde, formaldehyde, dimethyl suberimidate, carbodiimides, succinimidyls, diisocyanates, acyl azide, reuterin and/or genipin, its analog and/or its derivatives. The reversible crossliniking agent is selected from polyphenolic compounds, proanthocyanidin, epigallocatechin gallate, epicatechin, epigallocatechin and/or epicatechin gallate. The system is crosslinked by exposing the material to ultraviolet irradiation, dehydrothermal treatment,

107 M tris(hydroxymethyl)phosphine, ascorbate-copper, glucose-lysine or photo-oxidizers.

TECHNOLOGY FOCUS - POLYMERS - Preferred Polymer: The layer or zone is made of biodegradable shape memory polymer.

L150 ANSWER 7 OF 21 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN

AN 2005-283506 [29] WPIX

DNN N2005-232449 DNC C2005-088048

TI Responsive polymeric system, useful e.g. as a sealant, a transient barrier for the prevention of post-surgical adhesions and in the field of gene therapy, comprises one or more silicon-containing reactive groups.

DC A96 B04 D16 P32

IN COHN, D; SOSNIK, A

PA (YISS) YISSUM RES DEV CO HEBREW UNIV JERUSALEM

CYC 1

PI US 2005069573 A1 20050331 (200529)* 19 A61F002-00

ADT US 2005069573 A1 US 2004-845476 20040512

PRAI IL 2003-155866 20030512

IC ICM A61F002-00

AB US2005069573 A UPAB: 20050506

NOVELTY - Responsive polymeric system (I) comprises one or more silicon-containing reactive groups (where (I) undergoes a hydrolysis-condensation reaction primarily at a body site in the presence of water and at body temperature, and as a result of the reaction, the molecular weight of (I) increases due to polymerization and/or cross linking, and the rheological and mechanical properties of (I) are changed).

USE - (I) is useful as a sealant, a coating and lubricant, a transient barrier for the prevention of post-surgical adhesions, a matrix for the unimodal or multimodal controlled release of biologically active agents and in the area of tissue engineering and the field of gene therapy (claimed).

ADVANTAGE - (I) is biodegradable or selectively biodegradable. (I) is biodegradable where the system disappears from the site after a predetermined time. (All claimed.)

Dwg.0/9

FS CPI GMPI

FA AB; DCN

MC CPI: A06-A00B; A06-A00E; A06-A00E3; A07-A01; A07-A03; A07-A04F; A12-R08; A12-V03; A12-W11L; A12-W12; B04-C02; B04-C03; B04-E01; B04-F01; B04-N02; B11-C13; B12-M10A4; D05-H10; D05-H19

TECH UPTX: 20050506
TECHNOLOGY FOCUS - POLYMERS -

TECHNOLOGY FOCUS - POLYMERS - Preferred Components: (I) is deployable at the body site via a non-invasive or a minimally invasive surgical procedure. (I) comprises one or more alkoxysilane groups, which undergo a hydrolysis-condensation reaction in the presence of water, which reaction is effected primarily at a predetermined body site, the reaction resulting in an increase in the molecular weight of the polymeric system and producing a change in the rheological and mechanical properties of the system. (I) comprises one or more silanol groups which undergo a hydrolysis-condensation reaction in presence of water at an appropriate pH, which reaction is effected primarily at a body site, the reaction resulting in an increase in the molecular weight of the polymeric system and producing a change in the rheological and mechanical properties of (I). (I) is selectively biodegradable where the system reverts to an essentially un-polymerized or non-cross linked state after a predetermined time. (I) comprises at least one silicon-containing reactive group, the group being a mono, di or tri-functional group. (I) generates a polymer (a linear polymer, a block polymer, a graft polymer, a comb polymer, a

star-like polymer and/or a cross linked polymer). (I) also comprises additional reactive groups (hydroxyl, carboxyl, thiol, amine, isocyanate, thioisocyanate and/or double bond-containing active groups). The increase in the molecular weight of (I) and the change in its rheological and mechanical properties is partial and the system is still able to retain some degree of flowability. (I) comprises more than one component that form covalent bonds between them or generate physical blends or interpenetrating and/or pseudointerpenetrating networks at the predetermined body site. (I) contains at least one biomolecule to be delivered into the body. (I) contains living cells or a material of tissue origin. (I) also comprises a solid component, which is a macro, micro or nano-sized material (a polymer, a ceramic material, a metal, a carbon and/or a biological material), where the solid component is a particle, a sphere, a capsule, a rod, a slab, a fiber, a mesh, a ribbon, a web, a non-woven structure, a fabric, an amorphous lattice structure, a filament wound structure, a honeycomb structure and/or a braided structure and where the solid component may be hollow and/or porous. The solid component possesses reactive moieties capable of reacting with the reactive groups present in (I). The solid component is a biodegradable material. The solid component is a ceramic material (tricalcium phosphate and/or hydroxyapatite). The solid component is of tissue source. The solid component comprises a material (elastin, a collagenous material, albumin, a fibrinous material, demineralized tissue and/or an acellular tissue matrix). The solid component contains at least one biomolecule to be delivered into the body. The solid component contains living cells. The solid component is chemically or physically bound to (I). (I) is a low molecular weight polymer capable of being deployed at a predetermined body site by minimally invasive procedures, the low molecular weight polymer being (polyoxyalkylene, polyester, polyurethane, polyamide, polycarbonate, polyanhydride, polyorthoesters, polyurea, polypeptide, polyalkylene, acrylic or methacrylic polymers and/or polysaccharide). (I) is also capable of undergoing a transition that results in a sharp increase in viscosity in response to a predetermined trigger at a predetermined body site, where the transition results in an increase in the viscosity of (I) by at least about 2 times. The predetermined trigger is temperature, where the increase in viscosity takes place as a result of heating from a lower temperature to body temperature. (I) comprises water or an aqueous-based solvent. (I) is a polyoxyalkylene polymer, a block copolymer comprising polyethylene oxide (PEO) and polypropylene oxide (PPO) (a diblock, a triblock or a multiblock, a segmented block copolymer comprising polyethylene oxide (PEO) or polypropylene oxide (PPO) chains), where the PEO and PPO chains are connected via a chain extender, a poly(alkylco-oxyalkylene) copolymer having the formula R-(OCH2CH)n-OH (where R is an hydrophobic monofunctional segment (poly(tetramethylene glycol), poly(caprolactone), poly(lacticacid) and/or poly(siloxane)), a poly-(alkyl-co-oxyalkylene) copolymer having the formula (-R1-(OCH2CH) n-O)pH (where R1 is a bifunctional or multifunctional hydrophobic segment, a poly(N-alkyl substituted acrylamide), preferably poly(N-isopropyl acrylamide) and/or cellulose or its derivatives. The responsive component is a segmented block copolymer comprising polyethylene oxide (PEO) and polypropylene oxide (PPO) chains, where the PEO and PPO chains are connected via a chain extender, where the chain extender comprises a component (phosgene, aliphatic or aromatic dicarboxylic acids or their acyl chlorides or anhydrides, cyanuric chloride, dicyclohexylcarbodiimide (DCC), hexamethylene diisocyanate (HDI), methylene bisphenyldiisocyanate (MDI) or other aliphatic or aromatic diisocyanates). The poly(N-alkyl substituted acrylamide) is a copolymer comprising alkoxysilane-containing vinyl monomers. (I) further comprises other polymers that are responsive to

other stimuli (temperature, pH, ionic strength, electric and magnetic fields, energy sources covering a broad range of wavelengths (ultraviolet, visible, infrared, microwave, ultrasound, electron beam or x-rays radiation), fluids and/or biological species). The additional component is capable of undergoing a transition as a result of an increase in temperature that results in a sharp increase in viscosity of at least about 2 times. The responsive component contains biologically or pharmacologically active molecule/s, to be delivered into the body following a unimodal or multimodal time dependent release kinetics, as the molecular weight of (I) as well as its rheological and mechanical properties change at the predetermined body site. The responsive polymeric system contains biologically or pharmacologically active molecule/s, where the active molecules are covalently bound to (I) via silicon-containing reactive groups present in (I). The silicon moieties serve as nuclei for the deposition or crystallization of various materials. The silicon moieties serve as nuclei for the deposition or crystallization of hydroxyapatite or other calcium phosphate derivatives for bone regeneration induction at a predetermined body site. (I) is a water solution or a gel comprising a molecule containing silicon-containing reactive groups and a natural and/or synthetic macromolecule containing functional groups capable of reacting with the silicon-containing reactive groups at a predetermined body site. (I) is a water solution or a gel comprising a molecule containing silicon-containing reactive groups and functional groups capable of reacting with the silicon-containing reactive groups at a predetermined body site. The macromolecule comprises polymer or oligomer (alginates, hyaluronic acid, chitosan, and cellulose and their derivatives, collagen, gelatin, agarose, oligoHEMA, polyacrylic acid, polyvinyl alcohol, polyethylene oxide, TMPO, peptides and/or proteins).

UPTX: 20050506 ABEX

US 2004-565438P

US 2004-585775P

EXAMPLE - Pluronic F127 (molecular weight 12600, 25.2 g) were dried at 120degreesC under vacuum for 2 hours. Then isocyanatopropyl (1.2 g) and dioctotin compound (0.1 g) were added to the reaction mixture and reacted at 80degreesC for one hour, under mechanical stirring (160 rpm) and a dry nitrogen atmosphere. The polymer produced was dissolved in chloroform (30 ml) and precipitated in petroleum ether 40-60 (400 ml). The mixture was worked up to give pluronic F127di-(3-isocyanatopropyl)triethoxysilane.

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ΑN
     2005-121248 [13]
                        WPIX
CR
     1998-286610 [25]; 2004-118078 [12]; 2005-372235 [38]; 2005-417021 [42];
     2005-424463 [43]; 2005-580451 [59]; 2005-603254 [62]; 2005-603255 [62]
DNN N2005-104627
                        DNC C2005-040279
TΙ
     Crosslinked biodegradable stent or implant, useful for treating vascular
     restenosis, comprises layers or zones of biological material having
     bioactive agents, and being crosslinked with means for crosslinking
     biological material.
DC
     A96 B04 D21 D22 P32
IN
     CHEN, M; SUNG, H; TU, H; TU, P Y
PA
     (CHEN-I) CHEN M; (SUNG-I) SUNG H; (TUHH-I) TU H; (TUPY-I) TU P Y
CYC
                    A1 20050127 (200513)*
PΙ
     US 2005019404
                                                45
                                                      A61F002-00
ADT
     US 2005019404 A1 CIP of US 2003-610391 20030630, Provisional US
     2003-518050P 20031107, Provisional US 2004-547935P 20040226, Provisional
     US 2004-565438P 20040426, Provisional US 2004-574501P 20040526,
     Provisional US 2004-585775P 20040706, US 2004-916170 20040811
PRAI US 2004-916170
                          20040811; US 2003-610391
                                                         20030630;
                          20031107; US 2004-547935P
     US 2003-518050P
                                                         20040226:
                          20040426; US 2004-574501P
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L150 ANSWER 8 OF 21 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN

20040706

20040526;

IC ICM A61F002-00 ICS A61K009-22

AB US2005019404 A UPAB: 20050928

NOVELTY - A crosslinked biodegradable stent or implant (I) comprises one or more layers or zones of biological material having one or more bioactive agents, and being crosslinked with a material for crosslinking the biological material.

ACTIVITY - Cytostatic; Vasotropic; Antiarteriosclerotic; Vulnerary. No supporting data is given.

MECHANISM OF ACTION - None given.

USE - (M1) is useful for treating a target tissue of a patient, which involves providing (I) made one or more layers or zones of biological material comprising one or more bioactive agents, crosslinking the biological material, and delivering (I) to the target tissue and releasing the bioactive agent for treating the target tissue, where the target tissue comprises atherosclerotic plaque or vulnerable plaque (claimed). (I) is useful for treating tumor tissue, for treating tissue injury due to angioplasty, for treating vascular restenosis and atherosclerosis, and in other therapeutic applications.

ADVANTAGE - (I) comprises several layers or zones, each layer or zone comprising its own specific biodegradation rate and its specific loading characteristics, where the loading characteristics include drug type, drug releasing rate and combination of one or more drugs. (I) is loaded with several bioactive agents that are configured suitable for slow drug release, enabling effective treatment by each of several drugs. (I) slowly releases the drug to the surrounding tissue or lumen of the bodily cavity.

DESCRIPTION OF DRAWING(S) - The figure shows a longitudinal view of a vascular stent coated with drug-containing collagen layers that are crosslinked with genipin.

vascular stent 1
stent strut 2

collagen layers 5,6,7

tissue contact surface region 8A

blood contact surface region 8B

Dwg.7/18

FS CPI GMPI

MC

FA AB; GI; DCN

CPI: A12-V01; A12-V03; B01-B02; B01-D02; B02-A; B02-D; B02-E; B02-H; B02-R; B02-S; B02-T; B03-F; B04-A10; B04-C02B2; B04-C02E1; B04-H06; B05-B01G; B05-B01P; B05-C05; B06-A01; B06-A02; B06-A03; B06-D09; B07-A02B; B07-B01; B09-B; B10-A04; B10-A17; B10-B02A; B10-C03; B10-C04E; B10-D01; B11-C04A; B12-M16; B14-D07C; B14-D10; B14-F01G; B14-F07; B14-H01; D08-A; D09-C01

TECH UPTX: 20050224

TECHNOLOGY FOCUS - BIOTECHNOLOGY - Preferred Stent: (I) comprises first layer or zone of a first biological material with a first bioactive agent and a second layer or zone of a second biological material with a second bioactive agent. (I) further comprises a third layer or zone of a third biological material with a third bioactive agent, where at least one of the first and second layer or zone is made of a biodegradable shape memory polymer. The bioactive agent is chosen from analgesics/antipyretics, antiasthamatics, antibiotics, antidepressants, antidiabetics, antifungal agents, antihypertensive agents, antiinflammatories, antineoplastics, antianxiety agents,

inflammatories, antineoplastics, antianxiety agents,
immunosuppressive agents, antimigraine agents, sedatives/hypnotics,
antipsychotic agents, antimanic agents, antiarrhythmics, antiarthritic
agents, antigout agents, anticoagulants, thrombolytic agents,
antifibrinolytic agents, antiplatelet agents, antibacterial agents,
antiviral agents, antimicrobials, anti-infectives and their combinations.
The bioactive agent comprises an angiogenesis factor or anti-angiogenesis

factor. The bioactive agent is chosen from actinomycin D, paclitaxel, vincristin, methotrexate, angiopeptin, batimastat, halofuginone, sirolimus, tacrolimus, everolimus, ABT-578, tranilast, dexamethasone, mycophenolic acid and their combinations. The bioactive agent is chosen from lavastatin, thromboxane A2 synthetase inhibitors, eicosapentanoic acid, ciprostene, trapidil, angiotensin convening enzyme inhibitors, aspirin, heparin and their combinations. The bioactive agent is chosen from allicin, ginseng extract, ginsenoside Rgl, flavone, ginkgo biloba extract, glycyrrhetinic acid, lipostabil, proanthocyanides and their combinations. The bioactive agent is chosen from ApoA-I Milano or recombinant ApoA-I Milano/phospholipid complexes. The bioactive agent is chosen from biological cells or endothelial progenitor cells. The bioactive agent is chosen from growth factor such as vascular endothelial growth factor, transforming growth factor-beta, insulin-like growth factor, platelet derived growth factor, fibroblast growth factor and their combinations. The biological material is chosen from collagen, gelatin, elastin, chitosan, NOCC, fibrin glue, biological sealant, chitosan-alginate complex and their combinations. The biological material is crosslinked with a crosslinking agent chosen from genipin, its analog, derivatives and their combinations, aglycon geniposidic acid, epoxy compounds, dialdehyde, starch, glutaraldehyde, formaldehyde, dimethyl suberimidate, carbodiimides, succinimidyls, diisocyanates, reuterin, and acyl azide. The biological material is crosslinked with a means for crosslinking the material, the means comprising exposing the material to ultraviolet irradiation, dehydrothermal treatment, tris(hydroxymethyl)phosphine, ascorbate-copper, glucose-lysine or photo-oxidizers. The biological material is crosslinked with a reversible crosslinking agent chosen from polyphenolic compounds, proanthocyanidin, epigallocatechin gallate, epicatechin, epigallocatechin, epicatechin gallate, and their combinations. (I) is configured a cylindrical shape that has a first circumference length before contacting water and a second circumference length after contacting water, where the second circumference length is at least 5% more than the first circumference length. (I) is configured a cylindrical shape, comprising a several open-ring stent members configured in a cylindrical manner. (I) is configured a cylindrical shape comprising at least one spiral film.

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L150 ANSWER 9 OF 21 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN
     2004-775504 [76]
                        WPIX
DNN
    N2004-610972
                        DNC C2004-271511
     Osteoinductive powder useful as bone implant material for bone repair and
TΤ
     replacement comprises demineralized bone matrix particles, calcium
     phosphate powder and optionally biocompatible cohesiveness agent.
DC
     A96 B04 B05 D22 E33 P32
ΙN
     EGAN, D; GILLES, D P L D; LEE, D D; ROSENBERG, A D; TOFIGHI, A N; GILLES
     DE PELICHY, L D
PA
     (EGAN-I) EGAN D; (GILL-I) GILLES D P L D; (LEED-I) LEE D D; (ROSE-I)
     ROSENBERG A D; (TOFI-I) TOFIGHI A N; (ETEX-N) ETEX CORP
CYC
PΙ
     WO 2004091435
                     A2 20041028 (200476)* EN
                                                69
                                                      A61F000-00
        RW: AT BE BG BW CH CY CZ DE DK EA EE ES FI FR GB GH GM GR HU IE IT KE
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            KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NA NI NO NZ
            OM PG PH PL PT RO RU SC SD SE SG SK SL SY TJ TM TN TR TT TZ UA UG
            US UZ VC VN YU ZA ZM ZW
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A1 20050421 (200528)

2003-462416P 20030411, US 2004-822540 20040412

US 2005084542

ADT

WO 2004091435 A2 WO 2004-US11182 20040412; US 2005084542 A1 Provisional US

A61K035-32

PRAI US 2003-462416P 20030411; US 2004-822540 20040412

IC ICM A61F000-00; A61K035-32

ICS A61K033-42

AB WO2004091435 A UPAB: 20041125

NOVELTY - An osteoinductive powder (P) comprises demineralized bone matrix (DBM) particles, a calcium phosphate powder (a) and optionally a biocompatible cohesiveness agent. (P) forms a formable, self-hardening, poorly crystalline apatitic (PCA) calcium phosphate paste when mixed with a liquid.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

- (1) a formable, self-hardening, PCA calcium phosphate paste comprising a powder component containing DBM particles, calcium phosphate powder and optionally a biocompatible cohesiveness agent, and a fluid sufficient to produce a cohesive formable paste. The paste retains its cohesiveness when introduced at an implant site in vivo and hardens to form PCA calcium phosphate having a compressive strength of greater than 1 MPa;
- (2) a bone implant material comprising a PCA calcium phosphate. The PCA calcium phosphate is formed by combining a powder component including DBM particles, a calcium phosphate powder containing an amorphous calcium phosphate and a second calcium phosphate source and a biocompatible cohesiveness agent, and a fluid. The second calcium phosphate source is an acidic calcium phosphate. The powder component and the liquid combine to produce a paste that hardens to form a PCA calcium phosphate having a compressive strength of $1-20\,\mathrm{MPa}$; and
- (3) assaying the amount of DBM particles, by weight, in a sample comprising DBM particles and a calcium phosphate powder involving adding hydrogen chloride to the sample, agitating the sample, then obtaining, drying and weighing the extracted pellet of DBM particles.

ACTIVITY - Osteopathic.

MECHANISM OF ACTION - None given.

USE - As bone implant material for bone repair and replacement (claimed).

ADVANTAGE - The osteogenic bone implant composition approximates the chemical composition of natural bone. The organic composition of these implant is osteoinductive despite the presence of inorganic component and is present in an amount to maximize the regenerative capabilities of implant without compromising its formability and mechanical strength. The formulation is self-hardening PCA calcium phosphate paste suitable for formable paste which retains its cohesiveness having overall Ca/P ratio of less than 1.67 (preferably 1.0 - 1.67), a compressive strength of 1 - 20 (preferably 2) MPa and are remolded to bone in vivo. Dwg.0/1

FS CPI GMPI

FA AB; DCN

MC CPI: A12-V02; B04-B03C; B04-C01; B04-C02A; B04-C02B; B04-C02C; B04-C02D;

B04-C03; B04-E01; B04-G01; B04-H06L; B04-N02; B05-A01B; B05-B02A3;

B14-N01; D08-A; E31-K05C; E31-K06

TECH UPTX: 20041125

TECHNOLOGY FOCUS - INORGANIC CHEMISTRY - Preferred Composition: (P) comprises (wt.%): DBM particles (1 - 60, preferably 15) and (a) (50 - 99). (a) comprises an amorphous calcium phosphate (a1) and a second calcium phosphate (a2). DBM has a particle size of less than about 850 (preferably 125 - 850 or 53 - 125, especially less than 125) microm. (P) further comprises at least one cohesiveness agent, biologically active agent or effervescent agent.

Preferred Components: (a2) is an acidic (e.g. dicalcium phosphate dihydrate (preferred), calcium metaphosphate, heptacalcium phosphate, tricalcium phosphate, calcium pyrophosphate dihydrate, poorly crystalline

hydroxyapatite, calcium pyrophosphate, or octacalcium phosphate) or a neutral calcium phosphate. (al) and (a2) have an average crystalline domain size of less than 100 nm. (a) is subjected to a high-energy milling process prior to mixing with DBM particles. The cohesiveness agent is present in about 1 - 20 (preferably less than 5, especially less than 1) wt.%. The effervescent agent (1 - 40 wt.%) is sodium bicarbonate, carbon dioxide, air, nitrogen, helium, oxygen or argon.

TECHNOLOGY FOCUS - POLYMERS - Preferred Components: The cohesiveness agent is a polymer selected from polysaccharides, nucleic acids, carbohydrates, proteins, polypeptides, poly(alpha-hydroxy acids), poly(lactones), poly(amino acids), poly(anhydrides), poly(ortho esters), poly(anhydride-co-imides), poly(orthocarbonates), poly(alpha-hydroxy alkanoates), poly(dioxanones), poly(phosphoesters), poly(L-lactide), poly(D,L-lactide), polyglycolide, poly(lactide-co-glycolide), poly(L-lactide-co-D, L-lactide), poly(D,L-lactide-co-trimethylene carbonate), polyhydroxybutyrate, poly(epsilon-caprolactone), poly(delta-valerolactone), poly(gamma-butyrolactone), poly(caprolactone), polyacrylic acid, polycarboxylic acid, poly(allylamine hydrochloride), poly(diallyldimethylammonium chloride), poly(ethyleneimine), polypropylene fumarate, polyvinyl alcohol, polyvinylpyrrolidone, polyethylene, polymethylmethacrylate, poly(ethylene glycol), poly(ethylene oxide), poly(vinyl alcohol), poly(vinylpyrrolidone), poly(ethyloxazoline), poly(ethylene oxide)-co-poly(propylene oxide) block copolymers, poly(ethylene terephthalate)polyamide, dextran (preferably alpha-cyclodextrin, beta-cyclodextrin, gamma-cyclodextrin or sodium dextran sulfate), alginic acid, arabic gum, guar gum, xanthan gum, gelatin, chitin, chitosan, chitosan acetate, chitosan lactate, chondroitin sulfate, N,Ocarboxymethyl chitosan, fibrin glue, hyaluronic acid, sodium hyaluronate, cellulose (preferably methylcellulose, carboxy methylcellulose, hydroxypropyl methylcellulose or hydroxyethyl cellulose), a proteoglycan, a starch (preferably hydroxyethyl starch or soluble starch), a pluronic, collagen, glycogen, a keratin and/or silk.

TECHNOLOGY FOCUS - ORGANIC CHEMISTRY - Preferred Components: The cohesiveness agent is carbon fibers, glycerol, a glucosamine, lactic acid or sodium glycerophosphate.

TECHNOLOGY FOCUS - BIOLOGY - Preferred Components: The biologically active agent is an antibody, an **antibiotic**, a polynucleotide, a **polypeptide**, a protein (preferably osteogenic protein e.g. BMP-2, BMP-4, BMP-5, BMP-6, BMP-7, BMP-10, BMP-12, BMP-13, and BMP-14), an anti-cancer agent (preferably alkylating agent, platinum agent, antimetabolite, topoisomerase inhibitor, antitumor **antibiotic**, antimitotic agent, aromatase inhibitor, thymidylate synthase inhibitor, DNA antagonist, farnesyltransferase inhibitor, pump inhibitor, histone **acetyltransferase** inhibitor, metalloproteinase inhibitor, ribonucleoside reductase inhibitor, TNF-alpha agonist, TNF-alpha antagonist, endothelin-A receptor antagonist, retinoic acid receptor agonist, immunomodulator, hormonal agent, anti hormonal agent, photodynamic agent, and tyrosine kinase inhibitor), a growth factor, or a vaccine.

ABEX UPTX: 20041125

EXAMPLE - An osteoinductive powder comprising fibrous bone matrix particles $(0.4~\rm g)$ and a calcium phosphate powder $(0.6~\rm g)$ was prepared. A formable, self-hardening paste was prepared by hydrating the powder $(1~\rm g)$ with physiological saline. The compressive strength of the paste was evaluated by loading the paste into cylindrical stainless steel molds

having 6 mm diameter and 12 mm height. The molds were then immersed into saline bath at 37 degrees C for 2 hours. The hardened sample was then removed from the molds and tested for compressive strength. The average compressive strength was measured as 12 + 1 MPa.

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L150 ANSWER 10 OF 21 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN
AN
     2004-431600 [40]
                       WPIX
     2001-514527 [56]; 2003-362994 [34]
CR
DNC
    C2004-161595
TΙ
    Composition, useful for the treatment of diseases characterized by the
    production of mucin e.g. steatorrhea, allergic inflammation and
     chronic obstructive pulmonary diseases, comprises biphenyl compounds.
DC
IN
     JONES, S; LEVITT, R C; MCLANE, M; NICOLAIDES, N C; ZHOU, Y
PΑ
     (GENA-N) GENAERA CORP
CYC
    107
PΙ
    WO 2004043392
                     A2 20040527 (200440)* EN
                                                83
                                                      A61K000-00
        RW: AT BE BG BW CH CY CZ DE DK EA EE ES FI FR GB GH GM GR HU IE IT KE
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         W: AE AG AL AM AT AU AZ BA BB BG BR BW BY BZ CA CH CN CO CR CU CZ DE
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            KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NI NO NZ OM
            PG PH PL PT RO RU SC SD SE SG SK SL SY TJ TM TN TR TT TZ UA UG UZ
            VC VN YU ZA ZM ZW
    AU 2003287621
                     A1 20040603 (200470)
                                                      A61K000-00
    EP 1562903
                     A2 20050817 (200554) EN
                                                      C07D213-02
         R: AL AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HU IE IT LI LT LU LV
            MC MK NL PT RO SE SI SK TR
ADT
    WO 2004043392 A2 WO 2003-US35864 20031110; AU 2003287621 A1 AU 2003-287621
     20031110; EP 1562903 A2 EP 2003-781868 20031110, WO 2003-US35864 20031110
FDT
    AU 2003287621 A1 Based on WO 2004043392; EP 1562903 A2 Based on WO
     2004043392
PRAI US 2002-290443
                          20021108
     ICM A61K000-00; C07D213-02
IC
     ICS A61K031-44; C07D401-02
     WO2004043392 A UPAB: 20050823
AB
    NOVELTY - Treatment of diseases characterized by the production of mucin
     comprises the administration of a composition (A) comprising at least one
     biphenyl compound.
          DETAILED DESCRIPTION - Treatment of diseases characterized by the
     production of mucin comprises the administration of a composition
     comprising at least one biphenyl compound of formula (I).
          X1-X9 = C, S, O or N; either
          R1-R11 = H, CF3, (substituted) alkyl, (substituted) aryl, halo, halo
     substituted alkyl, halo substituted aryl, cycloalkyl, hydroxyl, alkyl
     ether, aryl ether, amine, alkyl amine, aryl amine, alkyl ester, aryl
     ester, alkyl sulfonamide, aryl sulfonamide, thiol, alkyl thioether, aryl
     thioether, alkyl sulfone, aryl sulfone, alkyl sulfoxide, aryl sulfoxide or
     sulfonamide; or
          R1R2, R2R3, R3R4, R4R5, R6R7, R7R8 or R8R9 = a cycloalkyl ring or an
     (hetero)aryl ring;
          Y = C(0)R, H, carboxylate, alkyl carboxylate,
     sulfate, sulfonate, phosphate, phosphonate, amides of carboxylic
     acids, esters of carboxylic acids, amides of phosphoric acids,
     esters of phosphoric acids, amides of sulfonic acids, esters of sulfonic
     acids, amides of phosphonic acids, esters of phosphonic acids,
     sulfonamide, phosphonamide, tetrazole or hydroxamic acid;
          R = aryl, phosphonate, styryl, 3H-isobenzofuran-1-one-3-oxyl or
     3H-isobenzofuran-1-one-3-yl;
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R11, Y = a cyclic sulfonamide;

Z = O, N, S, C, sulfoxide or sulfone (when the atom is S, sulfoxide or sulfone, the groups R10 and R11 are not present and when the atom is N,

only R10 is present); dashed lines = single or double bond; m = 0-1; and n = 1-2. Provided that no more than one of X1-X9 is N. INDEPENDENT CLAIMS are also included for: (1) A therapeutic composition (B) formulated for inhalation delivery to the lungs, comprising at least one of talniflumate, flufenamic acid, niflumic acid, mefenamic acid and/or N-(3-fluorobenzyl)-3-aminoquinoline, their salts or prodrugs effective to decrease mucin production or mucin synthesis; and (2) A biphenyl compound of formula (II). X = S, N, O or CR;Y = CRR', O, NR6 CRR'-CRR1 or CR=CR; Z = NR5, O, S, CRR' or CRR'-CRR'; R1-R3 = H, 1-8C alkyl, 1-8C alkoxy, NH2, OH, halosubstituted alkyl or R4 = H, benzofuran-2-one compound of formula (1), benzoic acid compound of formula (2) or biphenyl compound of formula (3) (preferably H); Q = CR, NR6 or benzimidazole compound of formula (4); R5 = H or benzyl (preferably H);R6 = H, 1-8C alkyl, 1-8C alkoxy, OH or halo (preferably H); dashed lines = single or double bonds; and R, R' = H, 1-8C alkyl, 1-8C alkoxy, OH or halo. In radicals (2) - (3) attachment points shown are exemplary and are not defined in the specification. ACTIVITY - Antiasthmatic; Antiinflammatory; CNS-Gen.; Respiratory-Gen.; Antimicrobial; Gastrointestinal-Gen.; Antidiarrheic; Antiallergic. No biological data available. MECHANISM OF ACTION - Mucin synthesis inhibitor; Cyclooxygenase-2 inhibitor. USE - Compounds (I) are useful in the treatment of diseases characterized by the production of mucin (particularly acute/chronic sinusitis, asthma, (chronic) bronchitis, chronic obstructive pulmonary disease, an inflammatory lung disease, cystic fibrosis, an acute/chronic respiratory infectious disease, emphysema, gastrointestinal malabsorption syndrome, steatorrhea, diarrhea, allergic inflammation and bronchial hyperresponsiveness). The treatment of diseases with (I) reduces airway inflammation, epithelial related inflammation, inflammatory cells and gastrointestinal inflammation; down-regulates mediators (interleukin 9) of airway inflammation; decreases the number of goblet cells in the respiratory tract, epithelia and gastrointestinal tract and decreases the number of submucosal glands in the respiratory tract, sinuses and gastrointestinal tract (claimed). The biological effectiveness of (I) in treating bronchial hyperresponsiveness was tested. The compounds were able to suppress airway hyperresponsiveness and also over-production of mucus in the lung caused by exposure to antigens. ADVANTAGE - Compounds (I) are highly effective for the treatment of diseases characterized by the production of mucin. Dwg.0/23CPI AB; GI; DCN CPI: B04-C02B1; B04-C02E3; B05-B01E; B05-B01F; B05-B01N; B06-A02; B06-H; B07-H; B10-A08; B10-A10; B10-A18; B10-B04; B10-D03;

FS

FΑ

MC

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B10-E02; B10-E04; B10-F02; B10-G02; B10-H01; B10-H02; B14-A01;
          B14-A02; B14-C03; B14-D05C; B14-E02; B14-E10; B14-G02A; B14-K01;
          B14-L06; B14-L07; B14-N04
TECH
                    UPTX: 20040624
     TECHNOLOGY FOCUS - PHARMACEUTICALS - Preferred Compounds: Compounds (I)
     are bendroflumethiazide, analogues and derivatives of: anthranilic acid,
     2-aminonicotinic acid, 2-amino-phenylacetic acid or
     aminoquinolines and their salts or prodrugs (preferably talniflumate, a
     talniflumate derivative and their salts or prodrugs.
     Preferred Composition: Composition (A) or (B) is in the form of a liquid
     (aerosolized) or a powder. (A) further comprises expectorant
     (guaifenesin), mucolytic agent, antibiotic and/or decongestant
     agent. (A) further comprises stabilizing agent (cyclodextrin),
     absorption-enhancing agent (chitosan) and/or flavoring agent.
     (B) comprises talniflumate and/or a talniflumate derivative and their
     salts or prodrugs. (I) is administered as a prodrug (comprised of
     talniflumate).
     Preferred Method: Compounds (I) decrease mucin synthesis/secretion
     (chloride channel dependent) in cells that express an ICACC chloride
     channel (comprised by one or more calcium activated chloride channels)
     (particularly in the respiratory tract, upper respiratory tract,
     gastrointestinal tract and pancreas). (A) is administered via inhalation
     to the lungs or nasal passages. (I) also inhibits cyclooxygenase
     (specifically cyclooxygenase 2). (B) (particularly micronized composition)
     is formulated to increase the bioavailability of (I).
     For the treatment of chronic sinusitis, talniflumate is preferably used.
ABEX
                    UPTX: 20040624
     SPECIFIC COMPOUNDS - The use of Talniflumate (preferred); Flufenamic acid;
     Niflumic acid; Mefenamic acid; Bendroflumethiazide; and
     N-(3-fluorobenzyl)-3-aminoquinoline is specifically claimed as (I).
     15 compounds (II) are specifically claimed e.g. 5-(2-oxo-2-(2-(3-
     trifluoromethyl-phenylamino)-pyridin-3-yl)-ethyl)-5H-furo(3,4-b)pyridin-7-
     one (IIa).
     ADMINISTRATION - Administration of (I) is oral or via inhalation
     (claimed), at a dosage of 0.01-100 (preferably 0.1-10) mg/kg/day
     (systemically).
     DEFINITIONS - Preferred Definitions: In (I);
     Y = C(0)R or carboxylate;
     R = aryl, phosphonate, styryl, 3H-isobenzofuran-1-one-3-oxyl or
     3H-isobenzofuran-1-one-3-yl;
     R1-R11 = CF3 or alkyl;
    X6 = C \text{ or } N;
     n = 2;
     one Z group = NR10; and the other Z group is CR10R11 (where R10 is H and
     Rll is an amine group); and
     Y = sulfone (where Y and R11 form a cyclic sulfonamide).
L150 ANSWER 11 OF 21 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN
     2004-282793 [26]
ΑN
                        WPIX
     2004-282792 [26]
CR
DNN
    N2004-224240
                        DNC C2004-108629
ΤI
    Active bone and cartilage regenerating composition for induction of new
    bone and cartilage formation in mammals, comprises bone marrow cells and
     demineralized bone matrix or demineralized tooth matrix together with
     site-responsive polymer.
DC
    A25 A96 B04 B07 D22 P34
IN
    COHN, D; GUREVITCH, O; KURKALLI, B G S; SLAVIN, S; SOSNIK, A
PΑ
     (YISS) YISSUM RES DEV CO HEBREW UNIV JERUSALEM; (HADA-N) HADASIT MEDICAL
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RES SERVICES & DEV CO LT; (HADA-N) HADASIT MEDICAL RES SERVICES & DEV CYC 106 WO 2004022121 PΤ A1 20040318 (200426)* EN 99 A61L027-38 RW: AT BE BG CH CY CZ DE DK EA EE ES FI FR GB GH GM GR HU IE IT KE LS LU MC MW MZ NL OA PT RO SD SE SI SK SL SZ TR TZ UG ZM ZW W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NI NO NZ OM PG PH PL PT RO RU SC SD SE SG SK SL SY TJ TM TN TR TT TZ UA UG US UZ VC VN YU ZA ZM ZW AU 2003256055 A1 20040329 (200459) A61L027-38 EP 1585556 A1 20051019 (200569) EN A61L027-38 R: AL AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HU IE IT LI LT LU LV MC MK NL PT RO SE SI SK TR ADT WO 2004022121 A1 WO 2003-IL728 20030904; AU 2003256055 A1 AU 2003-256055 20030904; EP 1585556 A1 EP 2003-794037 20030904, WO 2003-IL728 20030904 FDT AU 2003256055 Al Based on WO 2004022121; EP 1585556 Al Based on WO 2004022121 PRAI WO 2002-IL736 20020904 ICM A61L027-38 ICS A61K035-28; A61K035-32; A61L027-26

AB

WO2004022121 A UPAB: 20051027

NOVELTY - An active bone and cartilage regenerating composition comprising bone marrow cells (BMC) and demineralized bone matrix (DBM) or demineralized tooth matrix together with a site-responsive polymer and optionally carrier, additive, diluent, and/or excipient, is new.

DETAILED DESCRIPTION - An INDEPENDENT CLAIM is also included for a kit for performing transplantation of BMC in admixture with DBM and a site-responsive polymer into any one of a joint, a cranio-facial-maxillary bone, an alveolar bone of maxilla and mandibula, spine, pelvis and long bones, or for construction or reconstruction of an extraskeletal bone, including for mechanical or biological support of artificial implants to the joint or of the joint or to the bone of a mammal, where the kit comprises DBM in powder, particle, string or slice form; a site-responsive polymer; a BM aspiration needle; an intra-osseous bone drilling burr; a needle with a thick lumen for infusion of viscous bone marrow-DBM-site-responsive polymer mixture; a 2-way lumen connector for simultaneous mixing of BMC with DBM and site-responsive polymer and diluent; a medium for maintaining BMC; optionally additional factors stimulating osteogenesis; and cryogenic mechanism for handling and maintaining BMC or BMC together with DBM.

ACTIVITY - Cytostatic; Osteopathic. No biological data given. MECHANISM OF ACTION - None given.

USE - The composition is used for induction of new bone and cartilage formation in mammals; transplantation of mesenchymal progenitor cells into any one of a joint, a cranio-facial-maxillary bone, an alveolar bone of maxilla and mandibula, spine, pelvis or long bones of a subject (i.e. a mammal, preferably a human); construction or reconstruction of an extraskeletal bone of a subject in need; restoring and/or enhancing the formation of a new hyaline cartilage and/or subchondral bone structure; mechanical or biological support of an artificial implant to a joint or of a joint or to a bone of a subject; treatment of a patient suffering from any one of hereditary or acquired bone disorder, hereditary or acquired cartilage disorder, a primary malignant bone or cartilage disorder, bone defects due to metastases or bone lesions due to a hematopoietic malignancy, particularly multiple myeloma, metabolic bone diseases, bone infections, conditions involving congenital or acquired bone or cartilage deformities and Paget's disease; treatment of a patient in need of any one of correction of complex fractures, bone replacement and formation of new bone in plastic or sexual surgery, by administering the composition into

the joint or bone (all claimed).

ADVANTAGE - The invention maintains integrity and shape of the transplanted complex, provides transplanted complex with the mechanical properties to temporarily meet the requirements of the organism (e.g. withstanding physical and mechanical pressure) throughout the period of tissue regeneration.

Dwg.0/10

FS CPI GMPI FA AB; DCN

MC CPI: A12-V02; B04-C02A; B04-F02; B04-H06L; B04-J01; B04-L01; B04-N02; B11-C04A; B14-H01; D09-C01D

TECH UPTX: 20040421

TECHNOLOGY FOCUS - POLYMERS - Preferred Component: The site-responsive polymer is polymeric system or biodegradable RTG polymer comprising silicon-containing reactive group(s). The composition comprises active agents, preferably bone morphogenetic proteins (BMPs), an immunosuppressant, an immunomodulator, an antibiotic, or anti-inflammatory agents. The RTG polymer comprises hydrophilic and hydrophobic segments covalently bound by chain extender(s) or coupling agent, having at least two functional groups. The hydrophilic and hydrophobic segments do not display Reverse Thermal Gelation behavior of their own at body temperature. The RTG polymer is a segmented block copolymer comprising polyethylene oxide (PEO) and polypropylene oxide (PPO) chains connected via a chain extender. The chain extender is a bifunctional, trifunctional or multifunctional molecule from phosgene, aliphatic or aromatic dicarboxylic acids, their reactive derivatives such as acyl chlorides and anhydrides, diamines, diols, aminoacids, oligopeptides, polypeptides, or cyanuric chloride or any other bifunctional, trifunctional or multifunctional coupling agent, or other molecules, synthetic or of biological origin, able to react with the mono, bi, tri, or multifunctional -OH, -SH, -COOH, -NH2, -CN or -NCO group terminated hydrophobic and hydrophilic components, and/or any other bifunctional or multifunctional segment. The polymer can be Pluronic, preferably Pluronic F127 or F108. It may a random (-PEG6000-O-CO-(CH2)4-CO-O-PPG3000-)n poly(ether-ester) or an alternating (-PEG6000-O-CO-O-PPG3000-)n poly (ether-carbonate). The silicon-containing reactive group is capable of undergoing a condensation reaction effected primarily at a body site in the presence of water and at body temperature. The reaction results in an increase in the molecular weight of the polymeric system due to polymerization and/or cross-linking and produces at least a partial change in the rheological and mechanical properties of the system. The responsive polymeric system comprises alkoxysilane group(s) capable of undergoing a hydrolysis-condensation reaction in the presence of water which reaction is effected primarily at a body site. The responsive polymeric system comprises silicon-containing reactive group(s). It generates a polymer from a linear polymer, a graft polymer, a comb polymer, a star-like polymer, and/or a cross-linked polymer. It also comprises additional reactive groups from hydroxyl, carboxyl, thiol, amine, isocyanate, thio-isocyanate, and/or double bond-containing active groups. It may comprise a solid component that is a biodegradable material or chemically or physically bound to the responsive polymeric system. It is a silicon-containing monomer, oligomer or low molecular weight polymer, from polyoxyalkylene, polyester, polyurethane, polyamide, polycarbonate, acrylic and methacrylic polymers, poly anhydride, polyorthoesters, polyurea, polypeptide, polyalkylene, and/or polysaccharide. It can also be polyoxyalkylene polymer, a block copolymer comprising polyethylene oxide and polypropylene oxide from a diblock, a triblock or a multiblock, a segmented block copolymer comprising polyethylene oxide and polypropylene oxide chains connected via a chain extender, a

poly(alkyl-co-oxyalkylene) copolymer having the formula R-(OCH2CH)n-OH, a poly(alkyl-co-oxyalkylene) copoly(siloxane) copolymer of formula (-R'-(OCH2CH)n-O)pH, poly(N-alkyl substituted acrylamide), preferably poly(N-isopropyl acrylamide), and/or cellulose or its derivative. The responsive polymeric system is a segmented block copolymer comprising polyethylene oxide and polypropylene oxide-chains, connected via a chain extender from phosgene, aliphatic or aromatic dicarboxylic acids, or their reactive derivatives such as acyl chlorides and anhydrides or other molecules able to react with the OH terminal groups of the PEO and PPO chains, such as dicyclohexylcarbodiimide (DCC), aliphatic or aromatic diisocyanates from hexamethylene diisocyanate or methylene bisphenyldiisocyanate or cyanuric chloride and/or any other bifunctional or multifunctional segment. The poly(N-alkyl substituted acrylamide) is a copolymer comprising alkosilane-containing vinyl monomers. The responsive polymeric system can be alginates and its derivatives, hyaluronic acid and its derivatives, collagen, gelatin, chitosan and its derivatives, agarose, cellulose and its derivatives, water soluble synthetic, semi-synthetic or natural oligomers and polymers from oligoHEMA, polyacrylic acid, polyvinyl alcohol, glycerol, polyethylene oxide, TMPO, oligo and polysaccharides, oligopeptides, peptides, proteins, enzymes, growth factors, hormones, and/or drugs. The DBM is of vertebrate origin or human origin.

R = hydrophobic mono-functional segment from watpoly(tetramethylene glycol), poly(caprolactone), and/or poly(lactic selecacid);
R' = bifunctional or multifunctional hydrophobic segment.
Preferred Property: The DMB has a particle size of 50-2500, preferably 250-500 micrometer.

Preferred Composition: The ratio between BMC and DBM is 1:1-20:1, preferably 4:1 by volume. The composition contains BMC-DBM mixture and RTG polymer at a ratio of 5:1-1:5 preferably 3:1-1:2. The number of bone marrow cells in the composition is 10 to the power of 6-4x10 to the power of 10 cells/ml.

ABEX

UPTX: 20040421

ADMINISTRATION - The composition can be administered directly on a joint bearing a damage in the osteo-chondral complex or in the cranium of an animal with a partial bone defect in the perietal bone.

EXAMPLE - Male Lewis rats were anesthetized by intraperitoneal injection of Ketamine. Microfracture drilling was inflicted in articular cartilage and subchondral bone in the intrachondylar region of the femur. The defect was filled with BMC (bone marrow cells) suspension mixed with DBM (demineralized bone matrix) powder with the supplement of RTG polymeric material. The skin was closed with stainless clips. The transplantation of the composition into performed full thickness damage in the osteo-chondral complex of the knee joint allowed to maintain smooth and uniform regenerating surface in the defect area for complete rehabilitation of the joint.

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L150 ANSWER 12 OF 21 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN
AN
     2004-141480 [14]
                        WPIX
     2004-131881 [13]
CR
DNC
    C2004-056417
     Treatment or prevention of microbial infection e.g. viral infections
     comprises administration of sulfated polysaccharide, e.g. sulfated
     dextran, with low percent of sulfation.
DC
     A11 A96 B04 C03
ΙN
     COMPER, W D
PΑ
     (COMP-I) COMPER W D
CYC 1
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- PΙ US 2003181416 A1 20030925 (200414)* A61K031-727 33 ADT US 2003181416 A1 Provisional US 2002-346629P 20020110, Provisional US 2002-366532P 20020325, Provisional US 2002-366533P 20020325, Provisional US 2002-402695P 20020813, US 2002-321756 20021217 PRAI US 2002-321756 20021217; US 2002-346629P 20020110; US 2002-366532P 20020325; US 2002-366533P 20020325; US 2002-402695P 20020813 ICICM A61K031-727 ICS A61K031-715 AΒ US2003181416 A UPAB: 20040226 NOVELTY - Treatment or prevention (P1) of a microbial infection involves administration of a composition comprising a sulfated polysaccharide (A1) having 6-13 (preferably 9-13) % of sulfur substitution per glucose residue
 - or a levorotatory sulfated polysaccharide having 6-20 % sulfur.

 DETAILED DESCRIPTION INDEPENDENT CLAIMS are included for the following:
 - (1) treatment or prevention (P2) of a microbial infection involving administering a periodate treated anionic polysaccharide, a co-charged anionic polysaccharide (A2), or a compound (A3) selected from cellulose sulfate, (14)-2-deoxy-2-sulfamido-3-0-sulfo-(14)- beta -D-glycopyranan (derivative of chitosan), 2-acetamido -2-deoxy-3-0-sulfo(14)- beta -D-glycopyranan (derivative of chitosan), Achranthese bidentata polysaccharide sulfate, Aurintricarboxylic acid, calcium spirulan, carboxymethylchitin, chemically degraded heparin (Org 31733), chondroitin polysulfate, copolymer of sulfonic acid and biphenyl disulfonic acid urea (MDL 10128), curdlan sulfate, cyanovirin-N (from cyanobacterium), fucoidin, galactan sulfate, glucosamine-6-sulfate (monosaccharide), glycyrrhizin sulfate, heparin, inositol hexasulfate, lentinan sulfate, mannan sulfate, N-acylated heparin conjugates, N-carboxymethylchitosan-N, O-sulfate, oligonucleotide-poly(L-lysine)-heparin complexes, pentosan polysulfate heparin, phosphorothioate oligodeoxynucleotides, polyacetal polysulfate, polyinosinic-polycytidylic acid, polysaccharides from indocalamus tesselatus (bamboo leaves), prunellin, Rhamnan sulfate,
 - (xylanopolyhydrogen sulfate), peptidoglycan DS-4152, periodate degraded heparin, phosphorothicate oligodeoxynucleotides, polyacetal polysulfate, polyinosinic-polycytidylic acid, polysaccharides from indocalamus tesselatus (bamboo leaves), prunellin, Rhamnan sulfate, ribofuranan sulfate, sodium lauryl sulfate, sulfate dodecyl laminarapentacside (alkyl oligosaccharide), sulfated bacterial glycosaminooglycan, sulfated dodecyl laminari-oligomer (alkyl oligosaccharide), sulfated ganglicsides, sulfated laminara-oligosaccharide glycosides synthesized from laminara-tetracse, laminara-pentacse, laminara-hexacse, sulfated N-deacetylatedchitin, sulfated octadecyl maltohexacside (alkyl oligosaccharide), sulfated octadecyl ribofumans, sulfated oligoxylan (heparin mimetic), sulfated xylogalactans, sulfatide (3' sulfogalactosylceramide), Sulfoeveman and xylomannan sulfate;
 - (2) a method (P3) of controlling the sulfation of sulfated polysaccharide administered in vivo to mammal involving providing the sulfated polysaccharide with a sulfation to eliminate or reduce binding of the polysaccharide by high charge density polyanion cell receptors and to provide anti-microbial activity to the sulfated polysaccharide, and administering the polysaccharide to a mammal;
 - (3) a pharmaceutical composition comprising a sulfated dextran having 6-13 % sulfur and a molecular weight of greater than 25000; and
 - (4) a prophylactic device (preferably condom) which is coated with a sulfated polysaccharide having 6-13 % sulfur.

ACTIVITY - Antimicrobial; Virucide; Antibacterial; Anti-HIV; Antiparasitic; Antiinflammatory; Antiarthritic.

An in vivo anti-viral activity of dextran sulfate and variants of sulfated dextrans was assessed in a pharmacokinetic study involving single

intravenous dose of 60 mg/kg commercially available dextran sulfate (approx. 17% sulfur) (DS) 40000 molecular weight and dextran sulfate (12.2% sulfate) (DES6) 40000 molecular weight given to three male and three female rats. Rats were Sprague-Dawley, previously cannulated in the vena cava. Blood was drawn at various times after injection and assessed for anti-HIV activity in an acute infectivity cytoprotection assay system utilizing HIV-1 RF virus with CNE-SS cells based on Witvrouw et al., J. Acqur. Immun. Def Syndr., 3:343-347, 1990. The results indicated that DS was highly toxic with only one rat surviving beyond 24 hours. In contrast good survival and circulating anti-HIV activity for as long as 120 hours after injection were observed in the DES6 treated rats. DES6 showed a prolonged half-life in the blood between 12 and 18 hours and an extended anti-viral activity beyond 72 hours.

MECHANISM OF ACTION - Microbial growth inhibitor. Human peripheral blood mononuclear cells blasted with phytohemagluttin (PHA) and interleukins-2 (IL-2) were counted. The cells were suspended in RPMI 1640 (RTM) (1 multiply 106 cells/ml) without phenol red supplemented with 15% fetal bovine serum, L-glutamine (2 mM), penicillin (100 U/ml), streptomycin (100 mu g/ml), gentamycin (10 mu g/ml) and IL-2 (20 U/ml). Fifty mu 1 of cells were then distributed to the inner 60 wells of a 96 well round bottom microtiter culture plate. Each plate contains cell control wells, virus control wells and experimental wells. Sulfated dextran (12.5% sulfur) was added to the microtiter plate followed by pretitered dilution of HIV-1 RoJo. The assay was incubated for 6 days in a humidified atmosphere at 37 deg. C, 5% CO2, after which supernatants were collected and IC50 value was determined, which was 1.6 mu g/ml.

USE - For treatment or prevention of a microbial infection ADVANTAGE - (A1)-(A3) have a degree of sulfation effective to enable maximal interaction of constituent sulfate groups with the microbe, which causes the infection. (A1)-(A3) are not endocytosed or degraded by cell receptor binding in the mammal and thus, retains antimicrobial activity in vivo. The compounds reduce or avoid the adverse effects e.g. toxicities associated with the oral or parenteral administration of conventional sulfated polysaccharides. The compounds are administered directly to the lymphatic system of a patient.

Dwg.0/5

FS CPÍ

MC

ro cri

FA AB; DCN

CPI: A03-A00A; A12-V01; B04-B03C; B04-C02; B04-C03D; B14-A01; B14-A02; B14-A04; B14-B02; B14-C03; B14-C09; C04-B03C; C04-C02; C04-C03D; C14-A01; C14-A02; C14-A04; C14-A06; C14-B02; C14-C03; C14-C09

TECH UPTX: 20040226

TECHNOLOGY FOCUS - PHARMACEUTICALS - Preferred Compound: When the molecular weight of (A1) is greater than 5000 (preferably 5000-1000000, more preferably above 25000, especially above 40000, particularly greater than 500000) g/mol, it cannot treat a herpes infection except dextrin sulfate, cyclodextrin or carrageenan.

Preferred Method: (P1) and (P2) further involves administration of an additional therapeutic agent

TECHNOLOGY FOCUS - ORGANIC CHEMISTRY - Preferred Compound: (A1) comprises D-glucopyranose residues linked by alpha-1,6 linkages or L-glucopyranose residues. (A2) is co-charged with carboxymethyl, sulfonate and/or sulfate groups.

ABEX

UPTX: 20040226

SPECIFIC COMPOUNDS - Sulfated dextran is specifically claimed as (A1).

ADMINISTRATION - The compounds are administered in a dosage of 0.001 - 200 (preferably 0.005 - 100) or 0.1 - 1500 mg/kg per day parenterally, orally, topically (claimed), mucosally (including sublingually, buccally,

rectally, nasally or vaginally) or transdermally. The parenteral administration includes subcutaneous, intramuscular, bolus injection, intraarterial and intravenous administration.

L150 ANSWER 13 OF 21 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN 2004-084489 [09] AN WPIX DNC C2004-034808 TΙ New cosmetic or dermatological compound, useful e.g. for treating dry

skin, dandruff or acne, comprising active agent bonded to polymer via spacer and released in presence of microbial enzyme.

DC A96 B05 C03 D21 E19

ΙN GHANDCHI, P; QUINN, F X

PΑ (OREA) L'OREAL SA

CYC 1

PΙ FR 2839451 A1 20031114 (200409)* 52 A61K031-085 <--

ADT FR 2839451 A1 FR 2002-5737 20020507

PRAI FR 2002-5737 20020507

IC ICM A61K031-085

> ICS A61K031-05; A61K031-122; A61K031-4412; A61P017-00

ICA C07C039-30; C07C049-717

AΒ 2839451 A UPAB: 20040205

NOVELTY - A new compound (I) comprises active agent(s) (A) having an active site (AS), a spacer (SP) and a polymer (PL), (A) being in inactive form when bonded to (PL) via (SP), where: (i) (A) is released from (SP), by cleavage at (AS), in presence of enzyme released by a microorganism; and (ii) (AS) is selected from about 50 specific groups, e.g. carboxylic or sulfonic acid, alcohol, amine, thiol or ester.

DETAILED DESCRIPTION - A new compound (I) comprises active agent(s) (A) having an active site (AS), a spacer (SP) and a polymer (PL), (A) being in inactive form when bonded to (PL) via (SP), where:

- (i) (A) is released from (SP), by cleavage at (AS), in presence of enzyme released by a mcroorganism; and
- (ii) (AS) is selected from alkene, diene, alkyne, alcohol, carboxylic acid, ester, anhydride, aldehyde, ketone, aldo-ketene, keto-ketene, ether, epoxide, peroxide, hemiacetal, acetal, amine, hydrazine, amide, imine, imide, hydroxylamine, hydroxamic acid, oxime, hydrazone, hydrazide, nitrile, isocyanate, azo, azidocarbonyl, nitro, nitrate, sulfenic acid, sulfinic acid, sulfonic acid, sulfate, sulfone, sulfoxide, thioacid, thioketone, thioester, thiol, thioether, disulfide, acid halide, ether halide, sulfenyl, halide sulfonyl halide or nitrite ion.

INDEPENDENT CLAIMS are included for:

- (a) the preparation of (I), by reacting (A) with (SP) and (PL);
- (b) cosmetic, dermatological or pharmaceutical compositions comprising (I) in a suitable medium; and
- (c) a cosmetic treatment method for the skin or hair, involving application of a (I)-containing composition.

ACTIVITY - Dermatological; antiseborrheic; antiinflammatory ; antialopecia; anorectic; antibacterial; fungicide.

MECHANISM OF ACTION - None given in the source material.

USE - The (I)-containing compositions are specifically used: (a) cosmetically for treating or preventing dry skin, loss of tone and/or elasticity of the skin, hyperpigmentation, skin aging symptoms, sensitive skin or wrinkles, for modulating microcirculation or regulating comedolytic activity, as slimming agents, for toning the hair and for treating dandruff; or (b) medicinally for treating inflammatory skin conditions, pathological whitening of the hair, severe skin diseases (e.g. xerosis), alopecia or acne or for modulating cutaneous energetic metabolism (all claimed).

ADVANTAGE - (A) in active form is released in controlled and selective manner in the region to be treated, specifically in an amount proportional to the quantity of enzyme released by a microorganism causing the disorder to be treated (e.g. Propionibacterium acnes or Pityrosporium ovale).

Dwg.0/0

FS CPI

FA AB; DCN

MC CPI: A10-E01; A12-V01; A12-V04A; A12-V04C; B04-C02A; B04-C02D; B04-C02E; B10-A03; B10-A14; B10-E04; B10-F02; B10-G02; B12-M10; B14-A01;

B14-A04; B14-C03; B14-E12; B14-N17; B14-R02; C04-C02A; C04-C02D; C04-C02E; C10-A03; C10-A14; C10-E04; C10-F02; C10-G02; C12-M10; C14-A01; C14-A04; C14-C03; C14-E12; C14-N17; C14-R02; D08-B09A;

E10-A03; E10-A14B; E10-E04; E10-F02; E10-G02

UPTX: 20040205

TECH

UPTX: 20040205
TECHNOLOGY FOCUS - PHARMACEUTICALS - Preferred Active Agents: (AS) is carboxylic acid, sulfonic acid, alcohol, amine, thiol, ether, ester, amide or nitrile. (A) are selected from anti-acne, antiseborrheic, antidandruff, antialopecia, antiinflammatory,

antidandruff, antialopecia, antiinflammatory, antiinflammatory, antiinflammatory, antioxidant, anti-free radical, antiwrinkle, antiaging, antiseptic, biocidal, depigmenting, pro-pigmenting, moisturizing, keratolytic, comedolytic, desquamating, sunscreen, skin coloring, dyeing, conditioning, slimming, microcirculation modulating or cutaneous energetic metabolism modulating agents or ceramides. More specifically (A) is selected from triclosan (or its hexanoate, tetradecanoate, octadec-9-enoate, 2-acetoxyoctanoate or 2-hydroxyoct-2-enoate ester), 1-hydroxy-2-pyrrolidone derivatives (especially piroctone 2-acetoxyoctanoate or piroctone octanoate), tropolone (or its 2-acetoxyoctanoate or octanoate), hinokitiol, octoxyglycerin or menthol.

TECHNOLOGY FOCUS - POLYMERS - Preferred Materials: (PL) has a solubility in water of at least 0.01% at 20degreesC under atmospheric pressure and is preferably a polysaccharide, especially guar gum, xanthan gum, pullulan (most preferred), carrageenate, agar, agarose, dextran, cellulose gel, alginate, chitin or chitosan. (SP) comprise a 1-50C (preferably 3-12C) linear, branched or cyclic, saturated or unsaturated carbon chain, containing at least two groups selected from amine, thio, carbamate, ether, ester, amide and CN. Preferably (SP) contains 1-5 heteroatoms selected from N, O, S and P. In particular (SP) is hexane-1,6-diisocyanate or octane-1,8-diisocyanate.

ABEX

ADMINISTRATION - (I) is specifically contained at 0.001-30 (preferably 0.05-20, especially 0.1-2) weight %, in an aqueous, aqueous alcoholic or oily solution, oil-in-water, water-in-oil or multiple emulsion, aqueous or oily gel, anhydrous liquid, paste or solid, spherule-based oil in water dispersion, white or colored cream, ointment, milk, lotion, serum, paste, foam or shampoo formulation (all claimed). (I) is optionally together with a wide range of specific additives or other active agents.

EXAMPLE - A solution of 10 g 5-chloro-2-(2,4-dichlorophenoxy)-phenol (triclosan) in 70 m dichloromethane was treated with 10 g triethylamine and 10.2 g phenol-blocked 6-isocyanato-hexanoyl chloride and stirred for 20 hours at room temperature. Work-up and chromatographic purification gave 5-chloro-2-(2,4-dichlorophenoxy)-phenyl phenol-blocked 6-isocyanato-hexanoate. A solution of 10 g of the product in 100 ml dimethyl sulfoxide (DMSO)was treated with a catalytic amount of dibutyl tin dilaurate, heated at 110degreesC for 6 hours, cooled and treated with petroleum ether. The precipitate was recovered, washed and dried to give 5-chloro-2-(2,4-dichlorophenoxy)-phenyl 6-isocyanato-hexanoate (II). A

solution of 0.107 g (II) in 46 ml DMSO was treated with a solution of 4.05 g pullulan in 54 ml DMSO, heated at 80degreesC under nitrogen for 8 hours, poured into ethanol and kept at 4degreesC overnight. The precipitated was filtered off, dialyzed against water and lyophilized to give pullulan modified with 1% (II), having formula (Ia). Tests showed that (Ia) was hydrolyzed by Propionibacterium acnes or Pityrosporium ovale culture supernatants to release triclosan. (Ia) was incorporated at 1 weight % in an anti-acne cream.

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L150 ANSWER 14 OF 21 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN
ΑN
     2004-061934 [06]
                        WPIX
CR
     2000-072517 [06]
DNN
    N2004-050215
                        DNC C2004-025257
TI
     Preparation of a magnetic-resonance imageable medical device comprises
     providing a coating of paramagnetic-ion/chelate complex on the medical
     device encapsulated by a first hydrogel.
DC
     A96 B04 B07 D22 P34 S01 S03 S05
ΙN
     JIANG, X; LI, J; STROTHER, C M; UNAL, O; YU, H
PΑ
     (WISC) WISCONSIN ALUMNI RES FOUND
CYC
    101
PΙ
     WO 2003094975
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                                                44
                                                      A61K049-08
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            MC MW MZ NL OA PT SD SE SI SK SL SZ TR TZ UG ZM ZW
         W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK
            DM DZ EC EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR
            KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ OM PH PL PT
            RO RU SD SE SG SK SL TJ TM TN TR TT TZ UA UG UZ VN YU ZA ZM ZW
     AU 2002353152
                     A1 20031111 (200442)
                                                      A61K049-08
                                                                      <--
     EP 1501552
                     A1 20050202 (200510)
                                          EN
                                                      A61K049-08
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            MK NL PT RO SE SI SK TR
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                     B2 20050524 (200535)
                                                      A61B005-055
     JP 2005525176
                     W 20050825 (200560)
                                                56
                                                      A61L029-00
ADT
    WO 2003094975 A1 WO 2002-US40007 20021213; AU 2002353152 A1 AU 2002-353152
     20021213; EP 1501552 A1 EP 2002-790131 20021213, WO 2002-US40007 20021213;
     US 6896874 B2 Provisional US 1998-86817P 19980526, Cont of US 1998-105033
     19980625, CIP of US 2002-96368 20020312, US 2002-142363 20020509; JP
     2005525176 W WO 2002-US40007 20021213, JP 2004-503058 20021213
    AU 2002353152 Al Based on WO 2003094975; EP 1501552 Al Based on WO
     2003094975; US 6896874 B2 Cont of US 6361759; JP 2005525176 W Based on WO
     2003094975
PRAI US 2002-142363
                          20020509; US 1998-86817P
                                                         19980526;
     US 1998-105033
                          19980625; US 2002-96368
                                                         20020312
IC
     ICM A61B005-055; A61K049-08; A61L029-00
         A61K049-18; A61L029-18; A61L031-00; A61L031-18; A61M025-00;
          A61M025-01; G01R033-28
AB
     WO2003094975 A UPAB: 20050920
     NOVELTY - Preparation (M1) of a magnetic-resonance imageable medical
     device (D1) involves providing a coating on the medical device in which a
     paramagnetic-metal ion/chelate complex (c1) is encapsulated by a first
     hydrogel. A chelate of the paramagnetic-metal-ion/chelate complex is
     linked to a functional group (f1). The functional group is an amine group
     or a carboxyl group.
          DETAILED DESCRIPTION - INDEPENDENT CLAIMS are included for:
          (1) A medical device (D2) capable of being magnetic-resonance imaged
     comprises a chelate linked to a functional group (f2) (preferably amino or
     carboxyl group), paramagnetic-metal ion coordinated with the
     chelate to form a paramagnetic-metal-ion/chelate complex (c2) and a first
     hydrogel encapsulating (c2);
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(2) Reduction (M2) of the mobility of paramagnetic-metal-ion/chelate

complexes (c3) covalently linked to a solid-base polymer (p1) of a medical device involving: a1) providing a medical device having (c3) covalently linked to (p1) of the medical device and a2) encapsulating at least one (c3) covalently linked to the medical device with a hydrogel (h1). (h1) reduces the mobility of (c3) and thus enhances the magnetic-resonance imageability of the medical device; and

(3) Manufacture (M3) of a magnetic-resonance-imageable medical device (D3) involving providing (D3) and cross-linking a chain (preferably polymer chain) with a first hydrogel to form a (h1) overcoat on at least a portion of (D3) (where the chain has a paramagnetic-metal-ion/chelate complex (c4) linked to it).

USE - The method is useful for coating a medical device (claimed) e.g. stents, coils, valves, catheter or biopsy needle in diagnostic MR; in performing an operation on a target or a device which itself is implanted in the body (human or animal) for therapeutic purpose e.g. stent or graft.

ADVANTAGE - The medical device is coated so that the devices are readily visualized, particularly, in T1 weighted magnetic resonance images. Because of the high signal caused by the coating, the entirety of the coated devices can be readily visualized during e.g. an endovascular procedure. The coating is capable of emitting magnetic resonance signals. The coating provides ability to emit magnetic resonance signals and permits visualization of the entirety of a device or instrument so coated as used in interventional MR procedures. The coating provides visibility in interoperative MR of surgical instruments after being coated with the signal-enhancing coatings; provides improved visualization of implanted devices so coated, e.g. stents, coils, and valves.

FS CPI EPI GMPI

FA AB; DCN

MC CPI: A10-E01; A11-B05; A12-V00V; B04-C01; B04-C02; B04-C03; B04-N02; B05-A03; B10-B01B; B11-C04; B11-C09; D09-C01

EPI: S01-E02A2; S03-E07A; S03-E09X; S05-D02B3

TECH

UPTX: 20040123

TECHNOLOGY FOCUS - INSTRUMENTATION AND TESTING - Preferred Device: At least a portion of (D1) and (D2) is made from (p1). (f2) is a functional group on (pl) is formed by ia) treating the substrate to yield (f2) on it. (D4) has a surface and the surface is at least partially made from or coated with (pl) including the polymer chain. Preferred Method: (M1) further involves i) treating (p1) to yield (f1) on it; ii) chill-setting the coating after the coating is provided on the medical device; iii) uses a cross-linker (preferably glutaraldehyde) to cross-link (pl) containing an amine group and the first hydrogel containing an amine group to form a hydrogel overcoat; iv) uses a cross-linker (preferably glutaraldehyde) to cross-link and first and second hydrogel to form a hydrogel overcoat. The cross-linker connects the amine group to an aldehyde moiety of the glutaraldehyde. (c1) and (c2) is covalently linked to the medical device. The polymer chain and (h1) are cross-linked using a cross-linker (preferably glutaraldehyde). (p2) and the polymer chain is not covalently linked to the (D1), (D2) and (D4). Step i) involves: (1) plasma treating (pl) with a plasma gas which is hydrazine, ammonia and/or a chemical moiety of a nitrogen-hydrogen combination to obtain a plasma-treating functional group which is an amine group; (2) plasma treating (p1) with a plasma gas which is carbon dioxide or oxygen to obtain functional group which is a carboxyl group; and (3) melt coating with a hydrophilic polymer or precoating with a hydrophilic polymer containing primary amine groups. The chelate is covalently linked to the functional group by an amide linkage. In (M1), (D1) and (M2), the linker or spacer molecule (preferably

lactam or diamine) links the chelate of (c1), (c2) and (c3) to (f1), (f2) and (f3) respectively. (c1) is mixed with the first hydrogel to produce

the coating. In (D2), the (p2), the first hydrogel and second hydrogel are cross-linked to produce a hydrogel overcoat using a cross-linker (preferably glutaraldehyde). ia) involves the step 1), and 3). al) further involves plasma treating at least one portion of (p1) of the medical device before covalently linking the complex to it to provide functional groups (f3) selected from amino and carboxyl groups of it; covalently linking (c3) to (f3). (c4) is linked to the chain by a functional group (f4) (preferably amine or carboxy group) or is covalently linked to (D4). (c4) is formed by coordinating a paramagnetic-metal-ion with the chelate. (f4) is formed by plasma treating (p1).

TECHNOLOGY FOCUS - POLYMERS - Preferred Components: Compound (p1) is polyethylene, polypropylene, polyester, polyamide, polytetrafluoroethylene, polyurethane, polyamino undecanoic acid, polydimethylsiloxane, polyglycol, polyoxyethylene, polysorbate 60, stearate and palmitate ester of sorbitol copolymerized with ethylene glycol, polyvinyl acetate phthalate, polyvinyl alcohol or polystyrene sulfonate. (f1) and (f2) is a functional group of a polymer (p2) or a second hydrogel. The polymer chain is a second hydrogel. The first hydrogel, the second hydrogel, and (h1) are selected from collagen, gelatin, hyaluronate, fibrin, alginate, agarose, chitosan, poly(acrylic acid), poly(acrylamide), poly(2-hydroxyethyl methacrylate), poly(N-isopropylacrylamide), poly(ethylene glycol)/poly(ethylene oxide), poly(ethylene oxide)-block-poly(lactic acid), poly(vinyl alcohol), polyphosphazene and/or polypeptide (preferably gelatin). (p2) and the polymer chain is poly(N(3-aminopropyl)methacrylamide) having repeating unit of formula -(CH2C(CH3)(C(O)-NH-CH2CH2CH2NH2)-. The polymer chain is poly(N(3-aminopropyl)) methacrylamide). (c4) is attached to the polymer chain by an amine group of poly(N(3-aminopropyl)methacrylamide). TECHNOLOGY FOCUS - INORGANIC CHEMISTRY - Preferred Components: The paramagnetic-metal ion of (M1), (D2), (M2) and (M3) is of formula M(n+). M = lanthanide (preferably gadolinium) or transition metal selected from iron, manganese, chromium, cobalt or nickel (preferably lanthanide); and n = at least 2.

ABEX

UPTX: 20040123

SPECIFIC COMPOUNDS - 10 Compounds are specifically claimed as the chelate e.g. diethylenetriaminepentaacetic acid.

EXAMPLE - A polyethylene sheet (4.5 in diameter and 1 mil thick) was placed in a capacitively coupled, 50 kHz, stainless steel plasma reactor and hydrazine plasma treatment of the polyethylene film was performed. The substrate film was placed on the lower electrode. First, the base pressure was established in the reactor. Then the hydrazine pressure was slowly raised by opening the valve to the liquid hydrazine reservoir. The following plasma conditions were used: base pressure = 60 mT; treatment hydrazine pressure = 350 mT; RF Power = 25 W; treatment time = 5 minutes; source temperature (hydrazine reservoir) = 60 degrees C; temperature of substrate = 40 degrees C. Surface atomic composition of untreated and plasma-treated surfaces were evaluated using XPS (X-ray photoelectron spectroscopy).

In a 25 ml dry flask, DTPA (diethylenetriaminepentaacetic acid) (21.5 mg) was added to anhydrous pyridine (8 ml). In a small vessel, 1,1'-carbonyldiimidazole (CDI) (8.9 mg), as a coupling catalyst, was dissolved in anhydrous pyridine (2 ml). The CDI solution was slowly added into the reaction flask while stirring and the mixture was stirred at room temperature (RT) for 2 hours. A hydrazine-plasma treated polyethylene film was then immersed in the resulting solution. The resulting mixture was purged with dry argon for 10 minutes. After reaction for 20 hours, the polyethylene film was carefully washed in sequence with pyridine,

chloroform, methanol and water. The surface was checked with XPS, and the results showed the presence of carboxyl groups demonstrated the presence of DTPA. GdCl3.6H20 (0.70 g) was dissolved in water (100 ml). The DTPA-treated polyethylene film was soaked in the solution for 12 hours. The film was then removed from the solution and washed with water. The surface was checked with XPS and showed two peaks at a binding energy (BE) of 153.4 eV and BE of 148.0 eV, corresponding to chelated Gd3+ and free Gd3+ respectively. The film was repeatedly washed with water until the free Gd3+ peak at 148.0 eV disappeared from the XPS spectrum. The relative surface atomic composition of untreated and treated polyethylene (PE) surfaces showed %Gd, %N, %O and %C for the untreated PE of 0, 0, 2.6 and 97.4; for hydrazine plasma treated PE of 0, 15.3, 14.5 and 70.2; for DTPA coated PE of 0, 5, 37.8 and 57.2 and for Gd coated PE of 1.1, 3.7, 35 and 60.3 respectively. MR signal enhancement was assessed by imaging coated sheets of polyethylene and polypropylene as prepared above with gradient-recalled echo (GRE) and spin-echo (SE) techniques on a clinical 1.5 T scanner. The sheets were held stationary in a beaker filled with a tissue-mimic, fat-free food-grade yogurt, and the contrast-enhancement of the coating was calculated by normalizing the signal near the sheet by the yogurt signal. The T1-weighed GRE and SE MR images showed signal enhancement near the coated polymer sheet. The T1, estimates near the coated surface and in the yogurt were 0.4 s and 1.1 s, respectively. No enhancement was observed near control sheets.

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L150 ANSWER 15 OF 21 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN
AN
     2004-011538 [01]
                        WPIX
DNN
    N2004-008510
                        DNC C2004-003245
    Method for providing sustained release of drug to moist tissue involves
TI
     applying to the moist tissue drug delivery device containing N,
    O-carboxymethyl chitosan to provide adherence to the
    moist tissue.
DC
    All Al4 A96 B05 B07 D22 P32
ΙN
    ELSON, C; KYDONIEUS, A
PΑ
     (CHIT-N) CHITOGENICS INC
CYC
    100
PΙ
    WO 2003082163
                    A1 20031009 (200401)* EN
                                                20
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    AU 2002252565
                                                      A61F013-00
                    A1 20031013 (200435)
                     A1 20050112 (200504)
     EP 1494633
                                          EN
                                                      A61F013-00
         R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT
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ADT
    WO 2003082163 A1 WO 2002-US10149 20020328; AU 2002252565 A1 AU 2002-252565
     20020328, WO 2002-US10149 20020328; EP 1494633 A1 EP 2002-721645 20020328,
    WO 2002-US10149 20020328
FDT
    AU 2002252565 Al Based on WO 2003082163; EP 1494633 Al Based on WO
     2003082163
PRAI WO 2002-US10149
                          20020328
     ICM A61F013-00
     ICS A61F002-00; A61F006-06
AB
    WO2003082163 A UPAB: 20040102
    NOVELTY - Providing sustained release of a drug to moist tissue involves
     applying to the moist tissue a drug delivery device including a level of
    N, O-carboxymethyl chitosan as a component to
    provide an adherence to the moist tissue. The drug delivery device further
```

contains a drug to be delivered to provide sustained release of the drug and permeation into the moist tissue or into the surrounding cavity. ACTIVITY - Antiinflammatory.

Six female rats were anesthetized using sodium pentobarbital (60 mg/kg) and subsequently sacrificed by cervical dislocation. Twelve femurs were harvested and stripped of connective tissue by sharp dissection. Excess connective tissue was removed from the rat femur by immersing the rat femurs in boiling water for thirty minutes. The femurs were then rinsed and air-dried. Each femur was immersed in 1 ml of 125I NOCC solution. The other half of the femur was used to manipulate the femur. Subsequently, the femur was either placed directly into a scintillation vial and then placed in a gamma -counter rack, or the femur was subjected to a uniform wash before placed into a scintillation vial and the gamma -counter rack. Four groups of three 125I NOCC treated femurs were subjected to either one wash, two washes, three washes, or no washes. A wash consisted of the uniform agitation of the femur in approximately 150 ml of PBS for five seconds. Two washes consisted of a wash, removing the femur from PBS for one second, and then repeating a wash. Hence, three washes consisted of a wash, removal of the femur, a wash, removal of the femur, and one last wash. The PBS solution was replaced for each group of femurs. The activity of 125I NOCC was evaluated by a Beckmangamma -counter. The amount of 125I NOCC adhered to a rat femur was calculated, which used the activity of 1 ml of 125I NOCC (7.2 multiply 107 CPM) and the activity of the 125I NOCC on the femur, (detected by the gamma -counter). The result indicated that 125I NOCC adheres to rat femur. After third wash, it was found that 9.5 multiply 10-3 micro 1/mm2 (or about 0.1 micro g NOCC/mm2) of 125I NOCC remained adhered to the rat femur.

MECHANISM OF ACTION - None given.

USE - For sustained release of a drug; for adhering moist tissue e.g. defective moist tissue including lung tissues, heart tissues and intestinal tissues, together; to deliver a systemic therapeutic effect; and for preventing surgical adhesions in moist tissue at the site of surgical incision in the treatment of mouth sores and periodontal disease (claimed).

ADVANTAGE - The method provides proper adherent system for use with moist tissue that can be tailored in terms of delivery time and compatibility through the use of additional structural material. Dwg.0/10

FS CPI GMPI

FΑ AB: DCN

MC

CPI: A10-E08C; A10-E23; A12-V01; A12-V02; B06-D01; B07-D04C; B08-C01; B10-A17; B11-C03; B11-C04; B11-C04A; B12-M10A; D09-C04B TECH UPTX: 20040102

TECHNOLOGY FOCUS - BIOLOGY - Preferred Tissue: The moist tissue comprises mucosal tissue (preferably oral cavity tissue, buccal tissue, vaginal tissue or ocular tissue) including gastrointestinal tract, serous cavity, pleural, pericardial, peritoneal cavities or bone tissues.

TECHNOLOGY FOCUS - PHARMACEUTICALS - Preferred Device: The device comprises a structural support material selected from rubber, plastic, resin, natural and/or synthetic polymers. The device further comprises a tissue sealant or a surgical adhesion barrier (preferably fibrin glue or a cyanoacrylate).

Preferred Drug: The drug is melatonin, chlorpheniramine, chlorhexidine and/or tetracycline. The drug can also be selected from beta-blockers, glaucoma treating drugs, progestins, estrogens, antifungal agent, antibacterial agents, anti-viral gents, proteins or peptides (preferably levonorgestrel).

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L150 ANSWER 16 OF 21 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN
     2002-241173 [29]
AN
                        WPIX
DNC
    C2002-072443
ΤI
    New immunogen composition comprising an antigen, a biocompatible polymer
     and a liquid vehicle, useful for e.g. stimulating an immune response both
     systemically and mucosally, or for altering reproductive cycle of the
    host.
DC
    A25 A96 B04 B07 D16
IN
    BLONDER, J P; COESHOTT, C M; RODELL, T C; ROSENTHAL, G J; SCHAUER, W H
PA
     (BLON-I) BLONDER J P; (COES-I) COESHOTT C M; (RODE-I) RODELL T C; (ROSE-I)
     ROSENTHAL G J; (SCHA-I) SCHAUER W H; (RXKI-N) RXKINETIX INC
CYC
    96
    WO 2001098206
                     A1 20011227 (200229)* EN
PΤ
                                                67
                                                      C01B025-00
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            DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ
            LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD
            SE SG SI SK SL TJ TM TR TT TZ UA UG UZ VN YU ZA ZW
                                                      A61K039-21
     US 2002025326
                     A1 20020228 (200229)
                                                                      <--
    AU 2001076831
                     A 20020102 (200230)
                                                      C01B025-00
    EP 1315672
                     A1 20030604 (200337)
                                           EN
                                                      C01B025-00
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            RO SE SI TR
     US 2004258702
                     A1 20041223 (200504)
                                                      A61K039-00
ADT
    WO 2001098206 A1 WO 2001-US20096 20010622; US 2002025326 A1 CIP of US
     2000-602654 20000622, Provisional US 2001-278267P 20010323, US 2001-888235
     20010622; AU 2001076831 A AU 2001-76831 20010622; EP 1315672 A1 EP
     2001-954595 20010622, WO 2001-US20096 20010622; US 2004258702 A1 CIP of US
     2000-602654 20000622, Provisional US 2001-278267P 20010323, CIP of US
     2001-888235 20010622, US 2004-828842 20040421
    AU 2001076831 A Based on WO 2001098206; EP 1315672 Al Based on WO
     2001098206
PRAI US 2001-278267P
                          20010323; US 2000-602654
                                                         20000622;
     US 2001-888235
                          20010622; US 2004-828842
                                                         20040421
TC.
     ICM A61K039-00; A61K039-21; C01B025-00
         A01N043-04; A61K009-06; A61K009-32;
         A61K009-38; A61K009-40; A61K009-52;
         A61K009-58; A61K009-64; A61K009-66;
         A61K031-715; A61K031-721; A61K031-722;
         A61K031-74; A61K039-02; A61K039-095;
         A61K039-10; A61K039-12; A61K039-245;
         A61K039-29; A61K039-38; A61K045-00;
          A61K047-00; A61K047-30; A61K051-12;
          C01B015-16; C01B023-00; C01B025-18
AB
    WO 200198206 A UPAB: 20020508
    NOVELTY - A new immunogen composition for stimulating an immune response
    when administered to a host, comprising an antigen, a biocompatible
    polymer and a liquid vehicle, where the polymer interacts with the liquid
     vehicle to impart reverse thermal viscosity behavior to the composition,
     so that the viscosity of the composition increases when the temperature of
     the composition increases at a temperature range.
          DETAILED DESCRIPTION - A new immunogen composition for stimulating an
     immune response when administered to a host, comprising an antigen, a
     biocompatible polymer and a liquid vehicle, where the polymer interacts
     with the liquid vehicle to impart reverse thermal viscosity behavior to
     the composition, so that the viscosity of the composition increases when
     the temperature of the composition increases at a temperature range. The
     composition further comprises an additive enhancing the immune response
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selected from a penetration enhancer and/or an adjuvant.

INDEPENDENT CLAIMS are also included for the following:

- (1) delivery vehicle compositions (DC1) comprising a drug in an amount to produce a desired biological response in a host, a reverse-thermal gelation biocompatible polymer, a liquid vehicle in which the polymer is at least partially soluble at some temperature, and an additive selected from a penetration enhancer and/or an adjuvant;
 - (2) methods of packaging and storing the immunogen compositions;
- (3) a method for delivering a drug to a host by administering a delivery vehicle composition comprising a drug, a reverse thermal gelation biocompatible polymer, liquid vehicle in which the polymer is at least partially soluble at some temperature, and an additive; and
- (4) a method for delivering an antigen to a host to stimulate an immune response.

ACTIVITY - Immunostimulant; Antibacterial.

The IgG antibody response to intranasal immunization at week 0 followed by booster immunization at weeks 1 and 3. The IgG anti-TT responses of mice immunized and boosted with TT in PBS, TT in F127/chitosan and TT in F127/LPC were compared. Results of these studies indicate that the animals treated i.n. with TT in PBS failed to generate a significant anti-TT immune response, with a geometric mean titer of 159.6. The group immunized with TT in F127/LPC had a measurable nti-TT IgG response with a geometric mean titer of 544. The group immunized with TT in F127/chitosan clearly demonstrated a significant systemic anti-TT IgG response with a geometric mean titer of more than 5000. Studies indicate that intranasal immunization with TT in F127/chitosan induces a significant systemic IgG anti-TT antibody response.

MECHANISM OF ACTION - None given in source material.

USE - The composition is useful for stimulating an immune response both systemically and mucosally, for delivering an antigen (or drugs) to a host to treat or prevent an infectious disease, for altering mammalian reproductive cycle, for reducing or eliminating degradation of the antigen and lowing for a relatively slow sustained administration of antigens to the host.

Dwg.0/12

FS CPI

FA AB; DCN

MC CPI: A12-W11L; B01-D01; B04-B04C1; B04-C02; B04-C03C; B04-F09C; B04-F10; B05-A01B; B10-A09B; B10-A22; B10-C02; B10-C04E; B12-M01; B14-A01; B14-G01; D05-H07; D05-H10

TECH

UPTX: 20020508 FECHNOLOGY FOCUS - BIOTECHNOLO

TECHNOLOGY FOCUS - BIOTECHNOLOGY - Preferred Composition: The temperature range is below 40 degrees C, particularly 1-37 degrees C. The composition is in the form of a flowable medium at least when the composition is at a first temperature (T1) and the composition is in a gel form at least when the composition is at a second temperature (T2), where T2 is higher than T1. T1 is 1-20 degrees C, and T2 is 25-37 degrees C. The polymer is substantially dissolved in the liquid vehicle when the immunogen composition is at T1, and at least a portion of the polymer comes out of solution in the liquid vehicle when the temperature of the composition is raised from T1 to T2.

Preferred Method: Packaging and storing a delivery vehicle composition comprises placing the composition in the form of a flowable medium, in a container and then raising the temperature of the composition to convert the composition to a gel form for storage. The gel form can be converted back to a flowable medium for administration to the host by lowering the temperature of the composition in the container. A drug can be administered to the host by administering a delivery vehicle composition, where the delivery vehicle composition comprises a drug, a reverse thermal gelation biocompatible polymer, liquid vehicle in which the polymer is at

least partially soluble at some temperature, and an additive consisting of a penetration enhancer and/or an adjuvant. Prior to the administration, the delivery vehicle composition is at a temperature lower than the physiologic temperature of the host, after the administration, and the host warms the delivery vehicle composition so that the temperature of the composition increases. The delivery vehicle composition is in the form of a flowable medium immediately prior to administration and the viscosity of the delivery vehicle composition increases after the administering when the temperature of the delivery vehicle composition increases. The delivery vehicle composition is in the form of a flowable medium immediately before administration and is converted to a gel form after administration. The drug delivery composition is administered to the host by placing the composition into an injection device for injection to the host. At least a portion of the drug delivery composition in the gel form adheres to a mucosal surface to retain the drug and the additive in the vicinity of the mucosal surface for delivery of the drug across the mucosal surface. The mucosal surface is selected from rectal, vaginal, ocular, oral, nasal, intestinal, pulmonary or aural mucosal surfaces. The drug comprises an antigen which stimulates an immune response, preferably a systemic immune response in the host. The immune response is a booster to a previous primary immunization of the host, where at least a portion of the delivery vehicle composition adheres to a mucosal surface within the host to retain the drug and the additive in the vicinity of the mucosal surface for delivery of the drug across the mucosal surface. The magnitude of the immune response is the same or greater than a comparison immune response generated by a comparison composition that is administered in the same way as the delivery vehicle composition. The comparison composition and the delivery vehicle composition are the same except for the absence of the polymer and/or the additive, where the polymer and/or additive are absent in the comparison composition. Delivering an antigen to a host to stimulate an immune response comprises introducing an immunogen composition into a host, and the immunogen comprises an antigen, a reverse thermal gelation biocompatible polymer, a liquid vehicle in which the polymer is at least partially soluble at some temperature, and an additive consisting of a penetration enhancer and/or an adjuvant for enhancing the immune response. The immunogen composition of the produces at least a humoral immune response in the host, which is preferably a human.

TECHNOLOGY FOCUS - POLYMERS - Preferred Polymer: The polymer is a polyoxyalkylene block copolymer comprising at least one block of a first polyoxyalkylene and at least one of second polyoxyalkylene. The first polyoxyalkylene is polyoxyethylene and the second polyoxyalkylene is polyoxypropylene. The polyoxyalkylene block copolymer has the formula HO(C2H4O)b(C3H6O)a(C2H4O)bH (I) or H(OCH2CH2)b(OC(CH3)HCH2)a(OCH2CH2)bOH (II).

For (I): a = 15-80; and b = 50-150. For (II): a = 20-80, and b = 15-60.

The (C2H4O)b blocks comprise at least 70% of the weight of the polyoxyalkylene block copolymer. The block copolymer comprises at least one block of a polyoxyalkylene, which is a polyoxypropylene or a polyoxyethylene. The first polyoxyalkylene is a polyoxyethylene and the second polyoxyalkylene is a polyoxypropylene. The polyoxyethylene comprises at least 70% weight of the polymer. The polyoxypropylene has the formula (C3H6O)b (III) or (OC(CH3)HCH2)b (IV):

b = an integer.

Preferred Composition: The antigen may also comprise at least one rotavirus

and at least one antigen derived from rotavirus, a polysaccharide, a peptide mimetic of a polysaccharide. The adjuvant comprises dimethyl dioctadecyl ammonium bromide (DDA), a CpG motif, a cytokine, or a chitosan material, preferably an N,O-carboxymethyl chitosan. The liquid vehicle comprises 60-85% weight of the composition, the antigen comprises 0.0001-5% weight of the composition, the polymer comprises 5-33% weight of the composition, and the additive 0.1-1.0% (or 0.01 -10 % in DC1) by weight of the composition. The additive comprises a penetration enhancer selected from poly-L-arginines, polyoxyethylenesorbitan, polyoxyethylene-9-lauryl.

TECHNOLOGY FOCUS - BIOLOGY - Preferred Composition: The antigen is derived from a bacterium, protozoa, fungus, hookworm, virus or their combinations. The antigen comprises a tetanus toxoid, a diphtheria toxoid, a non-pathogenic mutant of tetanus toxoid, a non-pathogenic mutant of diphtheria toxoid or their combinations. The antigen comprises at least one antigen from Bordatella pertussis, influenza virus, M. tuberculosis, a causative agent of childhood illness. The antigen may also comprise at least one rotavirus and at least one antigen derived from rotavirus, a polysaccharide, a peptide mimetic of a polysaccharide, or an antigen from Neisseria meningitiditis or Streptococcus pneumoniae, an Epstein-Barr virus, Hepatitis C virus (HCV), HIV or at least one antigen derived from Epstein-Barr virus, HCV or HIV. The antigen may further be selected from at least one molecule involved in a mammalian reproductive cycle, HCG, tumor-specific antigen, and an antigen from a blood-borne pathogen. The immunogen composition contains at least two antigens comprising: a first component consisting of tetanus toxoid, and/or a nonpathogenic mutant of tetanus toxoid; and a second component consisting of diphtheria toxoid, and/or a nonpathogenic mutant of diphtheria toxoid. The adjuvant of the composition comprises products of microorganisms, such as bacteria or yeast that can enhance uptake and presentation of antigens by antigen presenting cells. The additive also comprises a penetration enhancer selected from a bacterially-derived product, lysophosphatidylcholine, a CpG motif, a detoxified mutant of CT, a detoxified mutant of ET and an outer membrane protein of Neisseria meningitidis serogroup b

TECHNOLOGY FOCUS - ORGANIC CHEMISTRY - Preferred Additive: The additive comprises an adjuvant for the antigen, where the adjuvant is selected from a chitosan material, dimethyl dioctadecyl ammonium bromide (DDA), a CPG motif, and a cytokine. The additive comprises a penetration enhancer selected from a chitosan material, fatty acids, salts of fusidic acid, sodium lauryl sulfate, citric acid, salicylates, caprylic glycerides, capric glycerides, sodium caprylate, sodium caprate, sodium laurate, sodium glycyrrhetinate, dipotassium glycyrrhizinate, glycyrrhenitic acid hydrogen succinate, disodium salt, acylcarnitines or phospholipids. The additive comprises chitosan material, preferably at least one chitosan and a chitosan derivative, where the chitosan material comprises N,O- carboxymethyl chitosan.

TECHNOLOGY FOCUS - PHARMACEUTICALS - Preferred Formulation: The composition is in the form of disperse droplets in a mist produced by a nebulizer. The immunogen composition is contained within a nebulizer which can actuate to produce a mist comprising dispersed droplets of the composition. The nebulizer is a nasal nebulizer. The immunogen composition may also be contained within an injection device for administration by injection to the host.

ABEX UPTX: 20020508

ADMINISTRATION - The composition is administered through intranasal, sublingual, oral, subcutaneous, intramuscular, or intraperitoneal. The delivery vehicle composition (DC1) is in the form of dispersed droplets in a mist during the administration, where the mist is introduced into the nasal cavity of the host during. Administration comprises nebulizing the composition to form the mist (claimed). No dosage given in source material.

EXAMPLE - Tetanus toxoid (TT) solution was obtained containing 961 Lf TT per ml of solution and 1884 Lf TT per mg of protein nitrogen. Pluronic F127 stock solution was prepared at 30 or 34% (w/w) by dissolving the polymer in ice-cold phosphate buffer solution where complete dissolution was achieved by storing overnight at 4 degrees C. Chitosan stock solution was prepared at 3% (w/w) in a 0.9% (w/v) saline solution containing 0.1% (v/v) acetic acid, and heated overnight at 37 degrees C to dissolve the chitosan. The stock solutions were then mixed together to prepare formulations containing various combinations of 200 Lf/ml TT, 0.5% (w/w) chitosan, and 16.25% (w/w) Pluronic F127. Formulations were used to administer a dose of 1.5 Lf of TT per mouse.

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L150 ANSWER 17 OF 21 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN
     2001-616153 [71]
                        WPIX
DNC
    C2001-184407
ΤI
    Micro- and nano-capsules with cationic charges on surface are used in
     laundry and other detergents, skin cleansers, shampoos and skin and hair
DC
    A18 A28 A87 A96 A97 B07 D21 D25 E19 F06
ΙN
    EISFELD, W; KRUPP, U; BRAUN, V; LOSSACK, A; SCHEIDGEN, A
PA
     (HENK) HENKEL KGAA
CYC
    42
PΙ
    WO 2001062376
                    A1 20010830 (200171)* GE
                                                66
                                                      B01J013-02
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         W: AU BR CA CN CZ DZ HU ID IL IN JP KR MX PL RO RU SG SI SK UA US ZA
     DE 10008305
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                                                      A61K007-00
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                                          GE
                                                      B01J013-02
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ADT
    WO 2001062376 A1 WO 2001-EP1887 20010220; DE 10008305 A1 DE 2000-10008305
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     2000-10008307 20000223; AU 2001046459 A AU 2001-46459 20010220; EP 1257353
    A1 EP 2001-919315 20010220, WO 2001-EP1887 20010220; EP 1257353 B1 EP
     2001-919315 20010220, WO 2001-EP1887 20010220; DE 50104385 G DE
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     ES 2231467 T3 EP 2001-919315 20010220
    AU 2001046459 A Based on WO 2001062376; EP 1257353 A1 Based on WO
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PRAI DE 2000-10008307
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     ICM A61K007-00; B01J013-02; C11D017-00
         A61K007-02; A61K007-035; A61K007-48;
          A61K009-00; C11D001-645; D06M023-12
AB
     WO 200162376 A UPAB: 20011203
```

NOVELTY - Micro- and nano-capsules with cationic charges on the surface are claimed.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for:

- (a) laundry and other detergents containing usual contents and these capsules;
- (b) skin cosmetics and cleansers containing usual contents and these capsules;
- (c) shampoos and hair cosmetics containing usual contents and these capsules.

USE - The capsules are used in laundry and other detergents, skin cleansers and cosmetics and shampoos and hair cosmetics (all claimed). The detergents preferably are used for cleaning hard surfaces or washing dishes, textile treatments, especially universal and fine laundry detergents, pre-treatments, stain removers, after-treatments, e.g. softeners, upholstery and carpet cleaners (all claimed). The skin cleansers and cosmetics preferably are washing, shower and bath liquids and bars, body and face cremes and lotions, effervescent compositions, eye cosmetics and decorative cosmetics, e.g. lipstick, lip-gloss, make up, face powder, mascara, eyeliner, kohl, eye shadow, nail cosmetics etc.; and the hair cosmetics e.g. setting cremes, lotions and gels, hair sprays, pomades, rinses, cures, permanent waving agents, colors and bleaches (all claimed)

ADVANTAGE - Micro- and nano-capsules containing care components and perfumes are normally used in detergents and cosmetics that are rinsed off after treatment. However, the binding power between substrate surfaces (textiles, skin and hair) and micro- and nano-capsules is usually only slight. The present micro- and nano-capsules with cationic surface charges have very good substantivity towards substrates, especially textiles, skin or hair, so that at least a certain amount remains on the substrate, even after treatment with water.

Dwg.0/0

FS CPI

FA AB; DCN

CPI: A12-V04A; A12-V04C; A12-W05; A12-W12A; A12-W12B; B04-C02; B04-C03; B04-D01; B07-D09; B10-A22; B12-M11E; B14-R01; B14-R02; B14-R04; D08-B01; D08-B02; D08-B04; D08-B05; D08-B06; D08-B09A; D11-A02; D11-D01A; D11-D01B; E07-D09C; E10-A22; F03-J03

UPTX: 20011203

TECH

MC

TECHNOLOGY FOCUS - ORGANIC CHEMISTRY - Preferred Capsules: The capsules consist of a core enclosed in wall material; or they are compact or the cores contain no active components. They have a particle size of 10 nm to 1000 microm, preferably 1-60 microm, especially 1.5-40 microm, more especially 2-30 microm. The weight ratio of core to wall is 99.9:0.1 to 2:1, preferably 95:5 to 70:30, especially 90:10 to 75:25. The core material is released by thermal energy, pressure, a change in pH or osmosis, preferably on or after application to a surface, especially a textile. The surface consists (partly) of or is coated with a cationic polymer or cationic compound.

Preferred Cationic Compounds: Preferred cationic compounds are quaternary ammonium compounds (I), ester-quats (II) and (IV) and quaternary imidazolinium compounds (III) of the given formulae, short-chain, water-soluble quaternary ammonium compounds e.g.

trihydroxyethylmethylammonium methosulfate, alkyltrimethylammonium chlorides, dialkyldimethylammonium chloride and trialkylmethylammonium chlorides, protonized alkylamines, cationic quaternary sugar derivatives, alkylamidoamines and quaternary ester compounds:

R, R1 = acyclic 12-24C alkyl;

R2 = saturated 1-4C (hydroxy)alkyl;

R3 = R, R1, R2 or an aromatic group;

COR4, COR5 = aliphatic 12-22C acyl with 0-3 double bonds;

```
R6 = hydrogen (H) or hydroxyl (OH);

m, n, o = 1, 2 or 3;

X- = a halide, methosulfate, methophosphate or phosphate ion;

R9 = H or saturated 1-4 C alkyl;

R10 = aliphatic, (un)saturated 12-18 C alkyl or O(CO)R20;

R20 = aliphatic, (un)saturated 12-18 C alkyl;

R11 = aliphatic, (un)saturated 12-18 C alkyl;

Z = imino (NH) or oxygen (O);

X' = an anion;

q = 1-4;

R12 R13, R14 = 1-4 C alkyl, alkenyl or hydroxyalkyl;

R15, R16 = 8-28 C alkyl; and

r = 0-5.

Preferred Core Materials: Suitable core materials include perfumes, care
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Preferred Core Materials: Suitable core materials include perfumes, care components, vitamins and provitamins, e.g. vitamin A, vitamin C, vitamin E (alpha-tocopherol) and other tocopherols, vitamin F (polyene-fatty acids), panthenol (provitamin B5), beta-carotene (provitamin A) and its derivatives, plant extracts, antidandruff agents, ultraviolet filters, emollients (cosmetic oils), conditioners, glycerol, textile finishing agents, e.g. impregnants, finishes, conditioners, easy-care finishes, feel modifiers and soil release finishes, antistatic, antimicrobial agents and fungicides.

Preferred Compositions: The detergents contain textile cleaning components and capsules containing textile care components and preferably are in liquid to gel or solid form. The skin cleaners and cosmetics contain cleansing components and capsules with a core of refatting and/or cosmetic components. The shampoos and hair cosmetics preferably are combination products containing cleaning component(s) and capsules with cosmetic substances as capsule material.

TECHNOLOGY FOCUS - POLYMERS - Preferred Cationic Polymers: Preferred cationic polymers are quaternized protein hydrolyzates, polyquaternium-polymers, copolymers of polyvinylpyrrolidone (PVP) and dimethylaminomethacrylate, copolymers or vinylimidazole and vinylpyrrolidone, aminosilicone (co)polymers, polyquaternized polymers, cationic biopolymers based on chitin and cationic silicone oils. Preferred Capsule Materials: Suitable materials are natural and synthetic polymers, especially polymeric polysaccharides, e.g. agarose or cellulose, starch, chitin, chitosan, proteins, e.g. gelatin, gum arabic, (m) ethylcellulose, carboxymethylethylcellulose, hydroxyethylcellulose, cellulose acetates, polyamides, polycyanoacrylates, polylactides, polyglycolides, polyaniline, polypyrrole, polyvinylpyrrolidone, polystyrene, polyvinyl alcohol, polystyrene/maleic anhydride copolymers, epoxide resins, polyethylene-imines, styrene/methyl methacrylate copolymers, poly(meth)acrylates, polycarbonates, polyesters, silicones, mixtures of gelatin and water glass or polyphosphate, cellulose acetate and phthalate, gelatin and maleic anhydride/methyl vinyl ether copolymers, cellulose acetate-butyrate and any mixture of these, which can have cationic groups.

Preferred Core Materials: Suitable core materials include biopolymers, silicone oils, polymers for fixing hair and cationic polymers.

UPTX: 20011203

ABEX

EXAMPLE - Textile softening formulations contained 0.2, 0.5, 1.0, 2.0 or 5 weight% optionally cationically-modified microcapsules containing perfume (rose oil, peppermint oil and orange oil), which had various particle sizes, with medians at about 5, 20 and 40 microm. The ratio of core to wall was between 90:10 and 75:25 in all cases. The formulations also contained (A) 5.0, (B) 16.0, (C) 21.5 weight% cationic surfactant , (A, B, C) 0, (D) 3.0 weight% polyethylene dispersion and (A) 0.3, (B, C, D) 0.55 weight%

magnesium chloride hexahydrate, rest free perfume, dye, thickener, antifoam and water. Cotton textiles were treated with the softener solution for 5 minutes, using 10.3 g softener/kg dry washing and a goods:liquor ratio of 1:5. They were then spun, hung to dry for 1 day and kept in a cupboard for 2, 6 or 9 days or ironed after drying. In all cases, the perfume intensity was much stronger when cationically modified capsules were used.

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L150 ANSWER 18 OF 21 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN
AN
     2000-506528 [46]
                       WPTX
DNC C2000-152074
ΤI
     New N, O-substituted chitins and
     chitosans prepared by alkoxylation, N-substitution with aldoses or
     ketones followed by reduction, useful as additives for cosmetics, e.g.
     shampoos and suntan lotions.
DC
     A11 A96 B07 D21
IN
     RATHJENS, A; WACHTER, R
PΑ
     (COGN-N) COGNIS DEUT GMBH
CYC
PΙ
     DE 19857547
                     C1 20000803 (200046)*
                                                14
                                                      C08L005-08
    DE 19857547 C1 DE 1998-1057547 19981214
ADT
PRAI DE 1998-19857547
                          19981214
IC
     ICM C08L005-08
     ICS A61K007-00
AB
     DE 19857547 C UPAB: 20000921
     NOVELTY - A new N,O-substituted biopolymer is prepared
          (1) alkoxylation or acylation of chitins and/or
     chitosans using alkylene oxides and/or carboxylic acid
     anhydrides and/or acid chlorides;
          (2) N-substitution of the resulting intermediates with aldoses and/or
     ketones; and
          (3) reducing the resulting imino groups to amines.
          DETAILED DESCRIPTION - An N, O-substituted
     biopolymer is claimed, which is prepared by:
          (1) by alkoxylation or acylation of chitins
     and/or chitosans using alkylene oxides and/or carboxylic
     acid anhydrides and/or acid chlorides;
          (2) N-substitution of the resulting intermediates with aldoses and/or
     ketones; and finally
          (3) reducing the resulting imino groups to amines.
          An INDEPENDENT CLAIM is also included for preparation of the
     biopolymers as described above.
          ACTIVITY - Dermatological; antimicrobial.
          MECHANISM OF ACTION - None given.
          USE - The polymer is useful as a gel- or film-former, a moisture
     regulator and antimicrobial agent in cosmetics (claimed) e.g. in skin
     cleansers, moisturizers, hair sprays, shampoos, conditioners, shower gels,
     foam baths, suntan lotions and nail polishes.
          ADVANTAGE - The polymer has good solubility in alcohol and water,
     especially at a pH of 6 - 8. It is also compatible with anionic
     formulating ingredients. The derivatization with alkoxy or acyl
     and amine groups results in improved gel- and film-forming properties, as
     well as imparting both moisture regulating and antimicrobial effects.
     Dwg.0/0
     CPI
FS
FA
     AB; DCN
MC
     CPI: A03-A; A10-A; A10-E09; A12-V04; B04-C02E3; B14-A01;
          B14-N17; B14-R01; D08-B03; D08-B09A
TECH
                    UPTX: 20000921
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TECHNOLOGY FOCUS - POLYMERS - Preferred Polymer: The chitosans have an average molecular weight of 10000 - 5000000 and the degree of acetylation is 5 - 95 %. The reaction is preferably carried out using ethylene oxide and/or propylene oxide in stage (1), in which the molar ratio of chitin/chitosan (based on monomer units) to alkylene oxide is 1:0.1 - 1:10, and/or succinic acid anhydride, where the molar ratio of chitin/chitosan to anhydride is 1:0.1 - 1:1.5. The aldose or ketose is selected from glucose, mannose, galactose, maltose, lactose, cellobiose, melibiose, chitobiose diacetate and/or N-acetylglucosamine, and the molar ratio of chitin-chitosan to aldose/ketose is 10:1 -1:2. The imine is reduced using complex hydrides. Finished compositions may contain e.g. surfactants, oils, silicones, biogenic compounds (e.g. tocopherol or retinol), deodorants, anti-dandruff agents (e.g. climbazol) and UV filters. ABEX UPTX: 20000921 EXAMPLE - Propoxylated chitosan (7.6 q) with a 0.75 degree of substitution was dissolved with stirring in 0.2 M acetic acid (330 ml) at 20 degreesC. A solution of lactose (25.7 g) in water (225 ml) was slowly added and the mixture was stirred for a further 18 hours. Sodium cyanoborohydride (7.1 g) was added and after 24 hours' stirring, the resulting clear solution was neutralized with 50 weight % NaOH, precipitated with acetone (800 ml), washed with more acetone and dried to constant weight at 40 degreesC. An oil-in-water sun lotion was prepared, which contained the following ingredients (weight %): Cutina GMS (RTM) (4.0), Lanette O (RTM) (1.0), Plantaren 818 (RTM) (5.0), Finsolv TN (RTM) (2.0), dioctyl carbonate (6.0), Cetiol J 600 (RTM) (4.0), the substituted chitosan prepared as described above (1.0), panthenol/bisabolol (1.2), Copherol F1300 (RTM) (2.0), Neo Heliopan 303 (RTM) (10.0), Neo Heliopan AV (RTM) (2.0), Uvinul T 150 (RTM) (3.0), zinc oxide (5.0), 86 weight % glycerin (5.0) and water/preservatives (to 100). L150 ANSWER 19 OF 21 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN **2000-096810** [08] WPIX C2000-028047 DNC Preparation of microspheres used for delivery of bioactive agents, as a means of radio-imaging tissue and for delivery of agrochemicals. A18 A28 A32 A96 A97 B07 C07 AMSDEN, B G; LIGGINS, R T (ANGI-N) ANGIOTECH PHARM INC CYC A1 19991111 (200008) * EN 52 C08J003-14 RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL OA PT SD SE SL SZ UG ZW W: AE AL AM AT AU AZ BA BB BG BR BY CA CH CN CU CZ DE DK EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT UA UG US UZ VN YU ZA ZW AU 9935140 A 19991123 (200016) US 6224794 B1 20010501 (200126) B05D007-00 WO 9957176 A1 WO 1999-CA367 19990506; AU 9935140 A AU 1999-35140 19990506; US 6224794 B1 Provisional US 1998-84508P 19980506, US 1999-305857 19990505 AU 9935140 A Based on WO 9957176

ΑN

TI

DC

IN PΑ

PΤ

ADT

FDT

AB

PRAI US 1998-84508P

ICM B05D007-00; C08J003-14

9957176 A UPAB: 20000215

jan delaval - 17 november 2005

19990505

19980506; US 1999-305857

B01J002-06; B01J013-02; B01J013-04; B32B015-02; B32B017-02

NOVELTY - Preparation of microspheres comprises passing a first fluid composition comprising a polymer and a solvent through an orifice and directly into a second fluid composition comprising water and a microsphere-stabilizing agent, under one of two conditions.

DETAILED DESCRIPTION - Preparation of microspheres comprises passing a first fluid composition comprising a polymer and a solvent through an orifice and directly into a second fluid composition comprising water and a microsphere-stabilizing agent, under at least one of conditions (a) and

- the first composition flows through a first conduit along a (a) first path and exits the first conduit at the orifice, the second composition flows through a second conduit along a second path in an upstream to downstream direction, the first conduit is connected to the second and terminates at the orifice, the first and second paths being orientated at an angle between 0 and 180 deg. relative to each other;
- (b) the first composition being at a first temperature and the second composition at a second temperature wherein the boiling point of the solvent of the first composition is less than or near the second temperature; and forming a composition comprising water and microspheres, the microspheres comprising the polymer.

USE - The microspheres are suitable for use in the delivery of bioactive agents for animal aquarian and human use, as a means of radio-imaging tissue and for the controlled release of agrochemicals.

ADVANTAGE - The method provides a means for preparation of polymeric microspheres with control over the average particle size and particle size distribution.

Dwg.0/10

FS CPI

FΑ AB; DCN

MC CPI: A11-A04; A12-S09; B04-C03; B12-K04B; B12-M10; B12-M11E; C04-C03; C12-K04B; C12-M10; C12-M11E

UPTX: 20000215

TECH

TECHNOLOGY FOCUS - ORGANIC CHEMISTRY - Preferred Process: Using method (a), the first fluid composition is preferably injected through a needle into the second composition, wherein the second composition is flowing past the tip of the needle in an upstream to downstream direction herein the needle and path of the second composition are preferably at an angle of 45-90degrees relative to each other. This method provides a population of emulsion droplets, having an average volume diameter influenced by the surface tension at the interface of the needle tip and the second composition, by the velocity of the second composition flowing past the needle tip, by the viscosity of the second composition and/or by the diameter of the needle. For method (b), the first composition is at a first temperature lower than the boiling point of the solvent and the second composition at a second temperature greater than the boiling point of the solvent, or only slightly less than the b.pt. of the solvent . The solvent may be an organic solvent and the second composition may be located within a column having a top and a bottom, the second composition may be stirred at a controlled stirring rate and the first composition may be introduced to the second composition through the bottom of the column at a controlled introduction rate. The resulting product is a composition comprising water and microspheres, the microspheres comprising the polymer. The process may further comprise separating some or all of the solvent from some or all of the microspheres. The resulting microspheres have a volume average diameter of less than 300 microns, preferably between 50 and 150 microns. Preferred Polymers: The polymer may be a lipophilic polymer such as polyester (e.g. poly(lactide), poly(caprolactone), poly(glycolide), poly((-valerolactone) or copolymers thereof), poly(ethylene-co-vinyl acetate), poly(siloxane), poly(butyrolactone), poly(urethanes), hydrophilic polymers such as etheylene oxide and/or propylene oxide

polymers, carboxylated poly(ethylene), poly(phosphazene),

polysaccharides such as chitosan, N,O-carboxymethyl chitosan, O-carboxymethyl chitosan, N-carboxymethyl chitosan, alginate, methylcellulose, hydroxymethylcellulose, acacia or tragacanth, gelatin and proteins or polypeptides such as serum albumin and poly(amino acids) and blends, copolymers and combinations of these polymers. A preferred polymer is poly(lactide-co-glycolide) at a concentration in the solvent of 5-10 w/v%.

Preferred Stabilizing Agent: a preferred microsphere-stabilizing agent is selected from poly(vinyl alcohol), gum arabic, CARBOPOL, ethylated starches, carboxymethylcellulose, hydroxymethylcellulose and mixtures thereof. A preferred stabilizing agent is poly(vinyl alcohol) present at a concentration of 1-2w/v% in water.

Preferred Solvent: preferred solvents include dichloromethane, carbon tetrachloride, THF, EtOAc and polyethylene glcyol.

TECHNOLOGY FOCUS - POLYMERS - Microspheres are prepared by passing a first fluid composition comprising a polymer and a solvent through an orifice and directly into a second fluid composition comprising water and a microsphere-stabilizing agent under specified conditions. The polymer may be a lipophilic polymer such as polyester (e.g. poly(lactide), poly(caprolactone), poly(glycolide), poly((-valerolactone) or copolymers thereof), poly(ethylene-co-vinyl acetate), poly(siloxane), poly(butyrolactone), poly(urethanes), hydrophilic polymers such as etheylene oxide and/or propylene oxide polymers, carboxylated poly(ethylene), poly(phosphazene), polysaccharides such as chitosan, N, O-carboxymethyl chitosan, O-carboxymethyl chitosan, Ncarboxymethyl chitosan, alginate, methylcellulose, hydroxymethylcellulose, acacia or tragacanth, gelatin and proteins or polypeptides such as serum albumin and poly(amino acids) and blends, copolymers and combinations of these polymers. A preferred polymer is poly(lactide-co-glycolide). The stabilizing agent may also be polymeric in nature e.g. it may be selected from poly(vinyl alcohol), gum arabic, CARBOPOL, ethylated starches, carboxymethylcellulose, hydroxymethylcellulose and mixtures thereof. The solvent may be polyethylene glycol.

ABEX UPTX: 20000215

EXAMPLE - A 10% w/v PLGA (85:15 lactide:glycolide weight average M.W. 88,000) in dichloromethane solution was pumped at 2ml/min through a 31 gauge stainless steel, blunt-end needle. The needle was seated at a 90 degree angle within TEFLON-coated polyethylene tubing (internal diameter 0.25 inches), protruding through the tubing wall with the end of the needle approximately 1mm from the tube wall. A 1 w/v% PVA in reverse osmosis water solution was pumped at a constant rate of $16.5 \, \text{cm/sec}$ through the polyethylene tubing and past the needle. The interfacial tension was 71 dyne/cm and the viscosity of the PVA solution was $0.015 \, \text{g/(cm s)}$. the volume average diameter of the microspheres formed was $295 \, \text{microns}$ with a standard deviation of $13.8 \, \text{microns}$.

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L150 ANSWER 20 OF 21 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN
AN
     1997-535380 [49]
                        WPIX
     1999-131833 [11]
CR
DNN N1997-445781
                        DNC C1997-171087
ΤI
     Topical anti-hyperalgesic film-forming composition - useful for treating
     peripheral hyperalgesia and inhibiting post-injury pain..
     A96 B02 B03 B07 D22 P34
DC
     BALOGH, I; FARRAR, J J; KUMAR, V; MAYCOCK, A L; FARRAR, J; MAYCOCK, L
ΙN
PΑ
     (ADOL-N) ADOLOR CORP
CYC 71
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PΙ
    WO 9733634
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                                                      A61L025-00
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                     C 20041026 (200471)
                                           ΕN
                                                      A61L015-22
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                     T2 20050616 (200540)
                                                      A61K047-30
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    WO 9733634 A1 WO 1997-US3315 19970226; US 5667773 A US 1996-614027
     19960312; AU 9719847 A AU 1997-19847 19970226; EP 888141 A1 EP 1997-907990
     19970226, WO 1997-US3315 19970226; AU 715912 B AU 1997-19847 19970226; KR
     99071497 A WO 1997-US3315 19970226, KR 1998-703769 19980520; KR 297417 B
     WO 1997-US3315 19970226, KR 1998-703769 19980520; EP 888141 B1 EP
     1997-907990 19970226, WO 1997-US3315 19970226; DE 69729284 E DE
     1997-629284 19970226, EP 1997-907990 19970226, WO 1997-US3315 19970226; CA
     2223514 C CA 1997-2223514 19970226, WO 1997-US3315 19970226; DE 69729284
     T2 DE 1997-629284 19970226, EP 1997-907990 19970226, WO 1997-US3315
     19970226
FDT
    AU 9719847 A Based on WO 9733634; EP 888141 A1 Based on WO 9733634; AU
     715912 B Previous Publ. AU 9719847, Based on WO 9733634; KR 99071497 A
     Based on WO 9733634; KR 297417 B Previous Publ. KR 99071497, Based on WO
     9733634; EP 888141 B1 Based on WO 9733634; DE 69729284 E Based on EP
     888141, Based on WO 9733634; CA 2223514 C Based on WO 9733634; DE 69729284
     T2 Based on EP 888141, Based on WO 9733634
PRAI US 1996-614027
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REP
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IC
     ICM A61K031-00; A61K047-30; A61L015-22; A61L024-00;
          A61L025-00
         A61K007-40; A61K009-08; A61K047-38;
          A61L015-44; A61L026-00; A61P029-00
AΒ
          9733634 A UPAB: 20050624
     A topical anti-hyperalgesic composition for coating an injured or
     inflamed site is new. The composition comprises: (a) 1-65% of an
     anti-hyperalgesic compound incorporated in a film-forming polymeric
     material; (b) 1-76% of film-forming polymeric material which is capable of
     forming a continuous film at pH 5.5-8.5 and which contains O, N or S atoms
     in combination with Ca2+, Mg2+, Zn2+ or Ba2+ in a ratio in the range 7.7
     to 1; and (c) 23-34% of aqueous carrier.
          The film forming material is: (a) anionic carboxylated
     polysaccharides of an anionic carboxylated polysaccharide of
```

The film forming material is: (a) anionic carboxylated polysaccharides of an anionic carboxylated polysaccharide of pectin (D-galacturonoglycan), algin (anhydro-D-mannuronic acid and anhydro-L-guluronic acid residues), gum karaya (D-galacturonic acid, D-galactose or L-rhamnose); (b) anionic sulphonated synthetic polymer of polystyrene or polyaryl sulphone; and (c) cationic aminopolysaccharides of keratosulphate, chondroitin sulphate, hyaluronic sulphate, heparin, chitin or dermatan sulphate.

USE - The composition is useful for treating peripheral hyperalgesia and is useful for inhibiting post-injury pain associated with local inflammatory conditions including inflammation following infection, blisters, boils, acute skin injuries, abrasions, burns, cuts, contusions, surgical incisions, irritations, poison ivy, allergic rashes,

```
dermatitis, stings, bites and inflammation of joints.
          ADVANTAGE - The composition has no effect on the central nervous
     system.
     Dwg.0/0
     CPI GMPI
FS
FΑ
     AB; DCN
MC
     CPI: A12-V01; A12-V03A; B04-C03; B06-D06; B07-D05; B12-M02D; B14-C01;
          B14-C03; B14-G02A; B14-N17; D09-C04B
ABEQ US
          5667773 A UPAB: 19971211
     Topical anti-hyperalgesic film-forming composition, for coating an
     injured/inflamed site on a mammalian patient to reduce
     hyperalgesia at the site, comprises: (a) 1-65 wt.% of an antihyperalgesic
     compound, which is devoid of central nervous system side effects; (b) 1-76
     wt.% of a film forming polymeric material; and (c) 23-34 wt.% of an
     aqueous carrier. The film-forming material is capable of forming a
     continuous film at a pH of 5.5-8.5. The polymeric material has atoms
     (selected from N, O and S) containing polarisable
     electrons, in combination with a divalent cation (selected from Ca2+,
     Mg2+, Zn2+ and Ba2+). The ratio of the atoms containing the polarisable
     electrons to the divalent cations is 7.7 to 1. The film-forming material
     is selected from sodium ethylcellulose sulphate, sodium cellulose
     acetate sulphate, sodium carboxyethyl cellulose,
     chondroitin sulphate, dermatan sulphate, keratosulphate, hyaluronic acid,
     heparin, chitin, polyvinyl pyrrolidone, polyvinyl alcohol and
     polyethylene oxide.
          USE - The composition is useful in treating post-injury pain
     associated with local inflammatory conditions, including
     inflammation following infection, blisters, boils, acute skin
     injuries, abrasions, burns, cuts, contusions, surgical incisions,
     irritations from various sources, poison ivy, allergic rashes, dermatitis,
     stings, bites and inflammation of joints.
     Dwg.0/0
L150 ANSWER 21 OF 21 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN
     1995-006710 [01]
ΑN
                        WPIX
DNC
    C1995-002363
ΤI
     Complexes of iodine with chitosan or derivs. - prepared in absence
     of solvent, useful as topical disinfectants and cicatrising agents, or as
     deodorants in cosmetics..
DC
     A11 A96 B04 D21 D22
IN
     AFFAITATI, P; DE, ROSA A; ROSSI, A
     (BIOT-N) DEV BIOTECHNOLOGICAL PROCESSESS SNC; (IMSI-N) IMS INT MEDICAL
     SERVICE SRL; (DBPB-N) DBP DEV BIOTECHNOLOGICAL PROCESSES; (DPEL-I) DI
     PELLICCIA M T; (IMSI-N) IMS INT MEDICAL SERVICES SRL; (IMSI-N) IMS-INT
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                                                                      <--
ADT
    WO 9426788 Al WO 1994-IT52 19940428; EP 649437 Al EP 1994-916383 19940428,
     WO 1994-IT52 19940428; US 5538955 A WO 1994-IT52 19940428, US 1995-362568
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19950103; IT 1261474 B IT 1993-RM291 19930507; EP 649437 B1 EP 1994-916383
     19940428, WO 1994-IT52 19940428; DE 69414521 E DE 1994-614521 19940428, EP
     1994-916383 19940428, WO 1994-IT52 19940428; ES 2123790 T3 EP 1994-916383
     19940428; CA 2139509 C CA 1994-2139509 19940428, WO 1994-IT52 19940428
     EP 649437 Al Based on WO 9426788; US 5538955 A Based on WO 9426788; EP
     649437 B1 Based on WO 9426788; DE 69414521 E Based on EP 649437, Based on
     WO 9426788; ES 2123790 T3 Based on EP 649437; CA 2139509 C Based on WO
     9426788
PRAI IT 1993-RM291
                          19930507
REP 03Jnl.Ref; JP 04178329; US 4275194; US 5051256
IC
     ICM A61K000-00; C08B037-08
     ICS A01N059-12; A61K031-70; A61K031-73
AR
          9426788 A UPAB: 19950110
     Preparation of complexes (A) of iodine with chitosan (II) or its
     derivs.comprises a reaction in the absence of solvent; or by dissolving
     iodine in a non-ionic surfactant then the solution added to an aqueous
solution of
     (II) or absorbed onto (II) in powdered form that is solubilised in water.
     Complexes of formula (I) are also new: X(I2)n X = monomeric unit of
     chitin, chitosan (opt. N-carboxybutyl,
     acyl, or -carboxymethyl substd.) N,O
     -carboxymethylchitosan, N,O-chitosan
     sulphate or their salts; n = 0.01-15.
          USE - (I) is used as a disinfectant (e.g. for wounds, burns etc),
     cicatrising agent or deodorant, in pharmaceutical and cosmetic compsns.
     Iodine is the active antimicrobial while (II) stimulates tissue
     regeneration and wound healing.
          ADVANTAGE - (A) may contain >60% iodine in a form resistant to
     sublimation, and if iodine content at most50 weight %, they are soluble in
     acidic aqueous solns. to give solns. that do not stain the skin and have good
     film-forming properties. (A) gradually release iodine over a long period
     so do not damage treated tissue. This preparation does not require large
     quantities of solvent and reaction times may be reduced.
     Dwq.0/0
FS
     CPI
FΑ
     AB; DCN
     CPI: A10-E04A; A10-E09; A12-V01; A12-V03C1; A12-V04; B04-C02E3;
MC
          B05-C07; B14-A01; B14-N17B; B14-R03; D08-B09; D09-A01C
          5538955 A UPAB: 19960905
     A process for the preparation of a charge transfer complex of iodine with
     chitosan or a derivative thereof wherein the chitosan or
     the derivative thereof and the iodine are made to react in the absence of
     solvent.
     Dwq.0/0
=> d his
     (FILE 'HOME' ENTERED AT 13:42:09 ON 17 NOV 2005)
                SET COST OFF
     FILE 'REGISTRY' ENTERED AT 13:42:15 ON 17 NOV 2005
                E CHITIN/CN
L1
              1 S E3
L2
            315 S 1398-61-4/CRN
                E CHITIN
           3200 S E3
L3
L4
            924 S L3 NOT E4-E13, E16
L5
            608 S L4 NOT L1, L2
L6
             90 S L5 NOT SQL/FA
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L7
             89 S L6 NOT DNA
L8
             11 S L2 AND N
L9
              5 S L8 AND C2H4O3
L10
              5 S L9 AND CARBOXYMETHYL ETHER
L11
              1 S 52519~63-8
L12
              8 S L1-L3 AND N AND O
L13
              4 S L12 NOT SQL/FA
     FILE 'HCAPLUS' ENTERED AT 13:50:01 ON 17 NOV 2005
L14
            375 S L11
L15
           8628 S L1
L16
             12 S L15 (L) N(L)O
                SEL AN 1 12
L17
              2 S L16 AND E1-E4
                SEL RN
     FILE 'REGISTRY' ENTERED AT 13:54:51 ON 17 NOV 2005
L18
             16 S E5-E20
L19
              2 S L18 AND L1-L3
L20
             14 S L18 NOT L19
L21
              1 S L20 AND C10H17NO8
     FILE 'HCAPLUS' ENTERED AT 13:57:06 ON 17 NOV 2005
L22
              6 S L21
                SEL AN 3-6
L23
              4 S L22 AND E21-E28
L24
              5 S L17, L23
L25
              1 S US20050214255/PN OR (US2004-810742? OR WO2005-US10103)/AP,PRN
                E ELSON C/AU
L26
            158 S E3-E8, E18, E19
                E KYDONIEUS A/AU
L27
            149 S E3-E10
                E HENDERSON S/AU
L28
             64 S E3, E10
                E HENDERSON SUE/AU
L29
              6 S E5, E9, E10
                E CHITOGEN/PA, CS
L30
             12 S E5-E12
L31
              4 S L26-L29 AND CHITIN
L32
              1 S L30 AND CHITIN
L33
              4 S L31, L32
              2 S L33 NOT (48 OR 61)/SC,SX
L34
L35
              2 S L33 NOT L34
                SEL RN L34
     FILE 'REGISTRY' ENTERED AT 14:04:20 ON 17 NOV 2005
L36
             18 S E1-E18
L37
              4 S 1404-00-8 OR 56124-62-0 OR 89-57-6 OR 23214-92-8
L38
              2 S L36 AND (CHITIN OR L1-L3)
L39
              4 S L36 AND CHITOSAN
     FILE 'HCAPLUS' ENTERED AT 14:06:45 ON 17 NOV 2005
L40
          23166 S L38, L39
L41
             10 S L26-L30 AND L40
L42
              6 S L41 NOT L33
L43
              5 S L42 NOT 44/SC
L44
          28096 S L2, L3
L45
              0 S L26-L30 AND L44 NOT L41,L33
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FILE 'REGISTRY' ENTERED AT 14:09:24 ON 17 NOV 2005

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L46
           1830 S CHITOSAN
     FILE 'HCAPLUS' ENTERED AT 14:09:32 ON 17 NOV 2005
L47
          19914 S L46
     FILE 'REGISTRY' ENTERED AT 14:09:45 ON 17 NOV 2005
L48
              1 S L39 AND 1/NC
L49
            894 S 9012-76-4/CRN
     FILE 'HCAPLUS' ENTERED AT 14:09:55 ON 17 NOV 2005
L50
           2267 S L49
L51
             21 S L26-L30 AND L47, L50
L52
             11 S L51 NOT L33, L41
L53
             18 S L34, L43, L52
L54
             18 S L53 AND L14-L17, L22-L35, L40-L45, L47, L50-L53
L55
             17 S L54 AND N O
L56
             18 S L54 AND ?CHITOSAN?
L57
              3 S L54 AND ?CHITIN?
L58
             18 S L54-L57
L59
             17 S L58 AND ?CARBOXY?
L60
             18 S L58, L59
                SEL RN 18
     FILE 'REGISTRY' ENTERED AT 14:14:16 ON 17 NOV 2005
L61
             21 S E19-E39
L62
              3 S 865532-59-8 OR 865533-35-3 OR 865533-54-6
L63
              2 S L61 AND C5H9NO4
L64
              1 S L63 AND CHITOSAN
L65
              2 S L61 AND C6H8O7
L66
              1 S L65 AND CHITOSAN
L67
              1 S SUCCINIC ACID/CN
L68
           6216 S 110-15-6/CRN
                E C4H4O3/MF
L69
             43 S E3 AND OC4/ES
L70
              6 S L69 AND 2 5 NOT (14C# OR 13C# OR 11C# OR (D OR T)/ELS)
L71
              5 S L70 NOT DIOL
L72
              1 S L71 NOT (LABELED OR 5 HYDROXY)
L73
           1881 S 108-30-5/CRN
L74
              3 S L2, L3 AND L68, L73
                E C4H2O3/MF
L75
             16 S E3 AND OC4/ES
L76
              3 S L75 AND 2 5 NOT (14C# OR 13C# OR 11C# OR (D OR T)/ELS OR LABE
              1 S 108-31-6
L77
L78
          24151 S 108-31-6/CRN
L79
              0 S L2, L3 AND L78
     FILE 'HCAPLUS' ENTERED AT 14:24:27 ON 17 NOV 2005
L80
              2 S L62, L64, L66
L81
             23 S L17, L24, L25, L34, L43, L60, L80
L82
             23 S L81 AND L14-L17, L22-L35, L40-L45, L47, L50-L60, L80-L81
L83
             18 S L82 AND N O
L84
             23 S L82 AND (?CHITOSAN? OR ?CHITIN? OR ?CARBOXY? OR ?ACYL?)
L85
              1 S L84 AND L37
L86
              6 S L84 AND (5 AMINOSALICYLIC ACID OR DOXORUBICIN OR ?PEPTIDE? OR
L87
             10 S L84 AND NOCC
L88
             23 S L84-L87
L89
              4 S L14 AND L37
L90
             34 S L14 AND (5 AMINOSALICYLIC ACID OR DOXORUBICIN OR ?PEPTIDE? OR
L91
             35 S L89, L90
L92
              0 S L22 AND L37
```

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1.93
              O S L22 AND (5 AMINOSALICYLIC ACID OR DOXORUBICIN OR ?PEPTIDE? OR
              41 S L22, L91
1.94
L95
              40 S L94 AND (PD<=20040325 OR PRD<=20040325 OR AD<=20040325)
              41 S L94, L95
L96
                 SEL AN 2 14 16 18 21 22 33 36 37
L97
              32 S L96 NOT E1-E18
     FILE 'REGISTRY' ENTERED AT 14:40:04 ON 17 NOV 2005
L98
               1 S DIVINYL SULFONE/CN
     FILE 'HCAPLUS' ENTERED AT 14:40:10 ON 17 NOV 2005
L99
           1071 S L98 OR DIVINYLSULFONE OR DIVINYLSULPHONE OR DIVINYL() (SULFON
L100
              21 S L99 AND L14, L15, L22, L40, L44, L47, L50
L101
              0 S L100 AND L97
L102
              33 S L25, L97
L103
              1 S L102 AND L99, L100
L104
              33 S L102,L103 AND L14-L17,L22-L35,L40-L45,L47,L50-L60,L80-L97,L99
                 SEL HIT RN
     FILE 'REGISTRY' ENTERED AT 14:42:36 ON 17 NOV 2005
L105
              31 S E19-E49
L106
              25 S L105 AND (CHITOSAN OR CHITIN OR L2 OR L49)
L107
               9 S L106 AND N
L108
               4 S L107 AND 1/NC
L109
               9 S L107, L108
L110
              8 S L109 NOT C16H36N
               6 S L105 NOT L106
L111
L112
               1 S L111 AND PMS/CI
L113
               9 S L110, L112
L114
               5 S L111 NOT L113
     FILE 'HCAPLUS' ENTERED AT 14:45:47 ON 17 NOV 2005
L115
              33 S L113 AND L104
     FILE 'REGISTRY' ENTERED AT 14:46:12 ON 17 NOV 2005
     FILE 'HCAPLUS' ENTERED AT 14:46:31 ON 17 NOV 2005
     FILE 'WPIX' ENTERED AT 14:47:05 ON 17 NOV 2005
L116
               0 S L25
                E CHITOSAN/CN
L117
               1 S E12
                 E CHITIN/CN
                 E N.O-/CN
L118
               2 S RA1YBP/DCN
L119
            5327 S (A61K031-722 OR C08B037-08 OR C08L005-08)/IPC OR (B04-C02E3 O
                 E CHITIN/DCN
                 E E3+ALL
L120
            1746 S E4, E6
L121
            2377 S E8
L122
           1745 S E10
L123
           11530 S L118-L122 OR (?CHITOSAN? OR ?CHITIN?)/BIX
L124
              71 S L123 AND N O/BI, ABEX
L125
              60 S L124 AND ?CARBOX?/BIX
L126
              15 S L125 AND ?ACYL?/BIX
L127
              29 S L125 AND ?ACET?/BIX
L128
              35 S L126, L127
L129
              24 S NOCC/BIX
L130
              57 S L128, L129
L131
              32 S L130 AND A61K/IPC
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25 S L130 NOT L131
L132
                SEL AN 2 3 6 7 11 12 13 16 19
L133
              9 S L132 AND E1-E9
                SEL AN L131 18 30 31
L134
             29 S L131 NOT E10-E12
L135
             38 S L133, L134
L136
             20 S L135 AND (5 AMINOSALICYLIC ACID OR DOXORUBICIN OR ?PEPTIDE? O
L137
             1 S L135 AND (AMINOSALICYL? OR AMINO SALICYL?)/BIX
L138
             20 S L136, L137
                E 5-AMINOSALICYLIC/CN
L139
              1 S E4
                E DOXORUBICIN/CN
L140
              9 S E3-E17
                E INTERFERON/CN
L141
             76 S E3-E94
                E VALRUBICIN/CN
L142
              1 S E3
                E MYTOMYCIN/CN
L143
              1 S E4
L144
             88 S L139-L143
                SEL SDCN
                EDIT /SDCN /DCN
L145
              0 S E1-E90 AND L135
L146
             11 S L135 AND P220/M0, M1, M2, M3, M4, M5, M6
L147
             23 S L138, L146
L148
             21 S L147 AND (?CHITOSAN? OR ?CHITIN?)/BIX
L149
             19 S L148 AND N O/BI, ABEX
L150
             21 S L148, L149
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FILE 'WPIX' ENTERED AT 15:29:30 ON 17 NOV 2005

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